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#### Remarks

#### The Invention

One embodiment of the invention (claims 60, 63, 66, 67, 69-71, and 73-75) provides a bisubstrate inhibitor of a protein kinase. The inhibitor comprises (1) a nucleotide or nucleotide analog moiety comprising a triphosphate, (2) a peptide moiety which is a substrate for said protein kinase and comprises a tyrosine residue, a 2-amino-3-(4-amino-phenyl)-propionic acid residue, a serine residue, a 2,3-diamino-propionic acid residue, a threonine residue, or a 2,3-diamino-butyric acid residue, and (3) a tether linked to the tyrosine residue via its phenolic oxygen, the 2-amino-3-(4-amino-phenyl)-propionic acid residue via its 3-amino nitrogen, the threonine residue via its hydroxyl oxygen, the 2,3-diamino-propionic acid residue via its 3-amino nitrogen, the threonine residue via its hydroxyl residue, or the 2,3-diamino-butyric acid residue via its 3-amino nitrogen and linked to the nucleotide or nucleotide analog moiety via the gamma phosphate of the triphosphate. The tether is greater than or equal to 4.9 Å as measured from a gamma phosphorus of the nucleotide or nucleotide analog moiety to a proton donor of the tether formed by the phenolic oxygen, the aniline nitrogen, the hydroxyl oxygen, or the 3-amino nitrogen.

Another embodiment of the invention (claims 1-15, 58, 72, and 76) provides a bisubstrate inhibitor of insulin receptor kinase. The inhibitor comprises (1) a nucleotide or nucleotide analog moiety comprising a triphosphate, (2) a peptide moiety which is a substrate for said protein kinase and comprises a tyrosine residue or a 2-amino-3-(4-amino-phenyl)-propionic acid residue, and (3) a tether linked to the tyrosine residue via its phenolic oxygen or to the 2-amino-3-(4-amino-phenyl)-propionic acid residue via its aniline nitrogen and linked to the nucleotide or nucleotide analog moiety via the gamma phosphate of the triphosphate. The tether is greater than or equal to 4.9 Å as measured from a gamma phosphorus of the nucleotide or nucleotide analog moiety to a proton donor of the tether formed by the phenolic oxygen or the aniline nitrogen. One particular embodiment of the invention (claim 15) provides a bisubstrate inhibitor of insulin receptor kinase which is compound 2.

#### Claim Amendments

Claim 5-7 were amended to remove the commas between the amino acid residues of the recited peptide as suggested in the Office Action. The current format with a space between three letter codes for amino acids is the standard format in the art. Claim 15 was amended to recite "a bisubstrate inhibitor of insulin receptor kinase." Claim 4 was amended to recite "the peptide comprises a 2-amino-3-(4-amino-phenyl)-propionic acid residue." Claim 66 was amended to recite "a nitrogen atom replaces a hydroxyl oxygen on the tyrosine." The claims have not been narrowed in scope by these amendments.

Claims 1 and 60 were amended to recite that the nucleotide or nucleotide analog moiety comprises a triphosphate. Support can be found *inter alia* at page 7, paragraph 27: "Suitable moieties include ATP, ATPγ-S, GTP, CTP, TTP, UTP, GTPγ-S, CTPγ-S, TTPγ-S, [and] UTPγ-S." The nucleotides and nucleotide analogs described contain a triphosphate. Claims 1 and 60 were also amended to recite that the peptide moiety comprises a specific amino acid residue (tyrosine or 2-amino-3-(4-amino-phenyl)-propionic acid for claim 1; tyrosine, 2-amino-3-(4-amino-phenyl)-propionic acid, serine, 2,3-diamino-propionic acid, threonine, or 2,3-diamino-butyric acid for claim 60). Support for "tyrosine, 2-amino-3-(4-amino-phenyl)-propionic acid, serine, 2,3-diamino-propionic acid, threonine, and 2,3-diamino-butyric acid" residues can be found *inter alia* at page 8, paragraph 29:

In order to make particular inhibitors with suitable tethers, the tyrosine residue of irktide is modified so that the phenolic oxygen is replaced with a nitrogen. Similarly, for the inhibitor of PKA, the serine residue is modified by substituting a nitrogen for the hydroxyl oxygen. Similarly, for threonine protein kinases, the hydroxyl oxygen can be replaced with a nitrogen.

Replacement of a phenolic oxygen atom of a tyrosine residue with a nitrogen atom results in 2-amino-3-(4-amino-phenyl)-propionic acid. The structures of 2-amino-3-(4-amino-phenyl)-propionic acid and tyrosine are detailed in Attachment 1. Additional support for 2-amino-3-(4-amino-phenyl)-propionic acid can be found in Figures 1A (compound 2) and 1C (the intermediate compound prior to addition of bromoacetic acid).

Replacement of a hydroxyl oxygen atom of a serine residue with a nitrogen atom results in 2,3-diamino-propionic acid. The structures of 2,3-diamino-propionic acid and serine are shown in Attachment 1. Additional support for 2,3-diamino-propionic acid can be found in Figure 4 (intermediate compound prior to addition of bromoacetic acid and compound 4).

Replacement of a hydroxyl oxygen atom of a threonine residue with a nitrogen atom results in 2,3-diamino-butyric acid. The structures of 2,3-diamino-butyric acid and threonine are shown in Attachment 1.

Claims 1 and 60 were further amended to recite that the tether is "linked to the tyrosine residue via its phenolic oxygen or to the 2-amino-3-(4-amino-phenyl)-propionic acid via its aniline nitrogen" (claims 1 and 60) or "linked to the serine residue via its hydroxyl oxygen, the 2,3-diamino-propionic acid residue via its 3-amino nitrogen, the threonine residue via its hydroxyl residue, or the 2,3-diamino-butyric acid residue via its 3-amino nitrogen" (claim 60). Support can be found in Figures 1A and 4. Claims 1 and 60 were also amended to recite that the tether is also "linked to the nucleotide or nucleotide analog via the gamma phosphate of the triphosphate. Support can be found, for example, in Figure 1A (compound 2) and Figure 4 (compound 4), both of which show a tether linked to a gamma phosphate.

Claims 1 and 60 were also amended to recite that the tether is greater than or equal to 4.9 Å measured from a gamma phosphorus of the nucleotide or nucleotide analog moiety to a proton donor of the tether formed by the phenolic oxygen, the aniline nitrogen (claims 1 and 60), or the hydroxyl oxygen, or the 3-amino nitrogen (claim 60)." Support can be found, for example, in Figures 1A and Figure 4, both of which show a tether linked to a proton donor.

No new matter is added by these claim amendments.

## The Rejection of Claims 1-15, 58, 60, 63, 66-67, 69-71, and 74-76 Under 35 U.S.C. § 112, Second Paragraph

Claims 1-15, 58, 60, 63, 66-67, 69-71, and 74-76 stand rejected under 35 U.S.C. § 112, second paragraph as indefinite. In particular the rejection asserts that the recitation "the tether is  $\geq$  4.9 Å measured from a gamma phosphorus of the nucleotide or nucleotide

analog moiety to the proton donor" is indefinite because the specification fails to identify how a skilled artisan would perform such a measurement. Office Action, page 3, second paragraph. Applicants respectfully traverse this rejection.

The specification teaches the use of Cambridge Soft's Chem3D computer program package to calculate the distance between the gamma phosphorus of the nucleotide or nucleotide analog moiety and a proton donor of the tether assuming an extended conformation of the acetyl linker. "Distance between the anilino nitrogen and the gamma phosphorus was calculated using Chem3D assuming an extended confirmation of the acetyl linker." Page 4, paragraph 17. Thus, the distance between the gamma phosphorus and the proton donor is calculated using a three-dimensional conformation that is extended. "Extended" means that the structure is relaxed to allow the atoms (*i.e.*, the gamma phosphorus and the proton donor) to be as far apart as possible within a covalent structure, assuming standard atomic radii for the covalent bonds and standard bond angles. The specification clearly teaches a skilled artisan how to calculate the distance between the gamma phosphate and the proton donor.

Claim 15 also stands rejected under 35 U.S.C. § 112, second paragraph as indefinite because the terms "said insulin receptor kinase" and "the bisubstrate inhibitor of insulin receptor kinase" allegedly had no antecedent basis. Claim 15 was amended to recite "a bisubstrate inhibitor of insulin receptor kinase" in the preamble thus establishing proper antecedent basis for the two terms.

The Office Action also asserts that claim 66 is allegedly unclear in the recitation of "a nitrogen atom replaces a hydroxyl oxygen on a tyrosine" because the claims from which claim 66 depend do not require a tyrosine. Claim 63, the claim from which claim 66 depends, was amended to recite that the peptide moiety of the bisubstrate inhibitor comprises "a tyrosine residue."

Withdrawal of this rejection is respectfully requested in view of the amendment.

#### The Rejection of Claim 15 Under 35 U.S.C. § 112, First Paragraph

Claim 15 stands rejected under 35 U.S.C. § 112, first paragraph as allegedly containing new matter. In particular the rejection asserts that the recitation of "[a] bisubstrate inhibitor of insulin kinase" is new matter. The preamble of claim 15 was

amended to recite "a bisubstrate inhibitor of insulin <u>receptor</u> kinase." Withdrawal of this rejection is respectfully requested.

## The Rejection of Claims 1-14, 58, 60, 63, 66-67, and 69-76 Under 35 U.S.C. § 112, First Paragraph

Claims 1-14, 58, 60, 63, 66-67, and 69-76 stand rejected under 35 U.S.C. § 112, first paragraph as allegedly failing to provide an adequate written description. In particular the rejection asserts that the application fails to disclose a representative number of species for the claimed genus of bisubstrate inhibitors of insulin receptor kinase or the claimed genus of bisubstrate inhibitors of protein kinases. Applicants respectfully traverse this rejection.

To satisfy the written description requirement for a claimed genus, the specification may describe a representative number of species (1) by actual reduction to practice, (2) by reduction to drawings, or (3) by disclosure of relevant identifying characteristics sufficient to show the applicant was in possession of the claimed genus. Manual of Patent Examining Procedure § 2163(II)(A)(3)(a)(ii). Relevant identifying characteristics can be, for example, structure or other physical and/or chemical properties, functional characteristics coupled with a known or disclosed correlation between function and structure, or a combination of such identifying characteristics. *Id.* A representative number of species is inversely related to the skill and knowledge in the art. *Id.* The specification need only describe in detail that which is new or not conventional. *Hybritech v. Monoclonal Antibodies*, 802 F.2d 1367, 231 U.S.P.Q. 81 (Fed. Cir. 1986).

Independent claim 1 and dependent claims 2-14, 58, 72, and 76 are directed to bisubstrate inhibitors of insulin receptor kinase. Independent claim 60 and dependent claims 63, 66-67, 69-71, and 74-75 are directed to bisubstrate inhibitors of protein kinases. The bisubstrate inhibitors of insulin receptor kinase and protein kinases comprise (1) a nucleotide or nucleotide analog moiety comprising a triphosphate, (2) a peptide moiety which is a substrate for the insulin receptor kinase or protein kinase, and (3) a tether. The peptide moiety of the bisubstrate inhibitors of insulin receptor kinase comprises a tyrosine residue or a 2-amino-3-(4-amino-phenyl)-propionic acid residue. The peptide moiety of the bisubstrate inhibitors of protein kinases comprises a tyrosine

residue, a 2-amino-3-(4-amino-phenyl)-propionic acid residue, a serine residue, a 2,3-diamino-propionic acid residue, a threonine residue, or a 2,3-diamino-butyric acid residue. The tether is linked to the tyrosine residue via its phenolic oxygen or to the 2-amino-3-(4-amino-phenyl)-propionic acid residue via its aniline nitrogen (claims 1 and 60). The proton donor also can be linked to the serine or threonine residues via their hydroxyl oxygen or to the 2,3-diamino-propionic acid or 2,3-diamino-butyric acid residues via their 3-amino nitrogen (claim 60). The tether also is linked to the nucleotide or nucleotide analog moiety via the gamma phosphate of the triphosphate. The tether is greater than or equal to 4.9 Å as measured from the gamma phosphorus of the nucleotide or nucleotide analog moiety to a proton donor formed by the phenolic oxygen, the aniline nitrogen, the hydroxyl oxygen, or the 3-amino nitrogen.

The Office Action asserts that the bisubstrate inhibitors of protein kinases (claims 60, 63, 66-67, 69-71, and 73-75) and the bisubstrate inhibitors of insulin receptor kinase (claims 1, 4, 8-14, 58, and 76) are unlimited with respect to the structures and positioning of the nucleotide or nucleotide analog moiety, peptide moiety and tether. More specifically, the Office Action alleges that the nucleotide or nucleotide analog moiety and peptide moiety can be linked in any manner.

Claims 1 and 60, as amended, recite identifying characteristics of the bisubstrate inhibitors of insulin receptor kinase and protein kinases, respectively. First, claims 1 and 60, as amended, recite a nucleotide or nucleotide analog moiety which comprises "a triphosphate." The triphosphate is a chemical and structural property of nucleotides or nucleotide analogs. Further structural properties are recited in claims 2, 3, 8, and 9. The recited nucleotide and nucleotide analog moieties are not unlimited with respect to structure.

Second, claims 1 and 60, as amended, recite a peptide moiety which comprises a specific amino acid residue. Claim 1, as amended, recites a peptide moiety which comprises a tyrosine residue or a 2-amino-3-(4-amino-phenyl)-propionic acid residue. Claim 60, as amended, recites a peptide moiety that comprises a tyrosine residue, a 2-amino-3-(4-amino-phenyl)-propionic acid residue, a serine residue, a 2,3-diamino-propionic acid residue, a threonine residue, or a 2,3,diamino-butyric acid residue. Further

structural properties are recited in claims 4-7, 10-14, and 69-71. Thus the peptide moiety is not unlimited with respect to structure.

Third, claims 1 and 60, as amended, recite identifying characteristics regarding the relationship between the nucleotide or nucleotide analog moiety and the tether. Claims 1 and 60, as amended, recite that the nucleotide or nucleotide analog moiety is linked to the tether "via the gamma phosphate of the triphosphate." Thus the positioning of the nucleotide or nucleotide analog moiety is not unlimited.

Identifying characteristics of the linkage between the peptide moiety and the tether are also recited. Claim 1, as amended, requires the tether to be linked to the tyrosine residue of the peptide moiety via its phenolic oxygen or to the 2-amino-3-(4-amino-phenyl)-propionic acid residue of the peptide moiety via its aniline nitrogen. Claim 60, as amended, requires the proton donor of the tether to be linked to the tyrosine residue of the peptide moiety via its phenolic oxygen, to the 2-amino-3-(4-amino-phenyl)-propionic acid residue of the peptide moiety via its aniline nitrogen, to the serine or threonine residues via their hydroxyl oxygen, or to the 2,3-diamino-propionic acid or 2,3-diamino-butyric acid residues via their 3-amino nitrogen. Thus the positioning of the peptide moiety is not unlimited.

The previous amendment (dated March 17, 2004) identified nineteen peptides that have been shown in the prior art to be natural substrates of insulin receptor kinase or to function as substrates for the insulin receptor kinase. All nineteen peptides contain a tyrosine residue. Using these nineteen peptides, the ten nucleotides and nucleotide analogs which comprise a triphosphate taught in the specification, a simple 2-carbon tether taught in the specification, and the linking requirements recited in claim 1, at least 190 bisubstrate inhibitors of insulin receptor kinase can be constructed (19 peptides x 10 nucleotides or nucleotide analogs x 1 tether = 190 bisubstrate inhibitors of insulin receptor kinase). Using a 2-amino-3-(4-amino-phenyl)-propionic acid residue for the tyrosine residue (as disclosed at page 8, paragraph 29) in each of the nineteen peptides an additional 190 bisubstrate inhibitors of insulin receptor kinase (19 peptides x 10 nucleotides or nucleotide analogs x 1 tether = 190 bisubstrate inhibitors of insulin receptor kinase) are disclosed. Thus, 190 bisubstrate inhibitors of insulin receptor kinase are disclosed. This is a representative number of species.

The peptide moiety of claim 60 "is a substrate for said protein kinase." Eightytwo peptides were identified in the previous amendment that were known in the prior art to be natural substrates of protein kinases or to function as substrates for the protein kinases. All 82 peptides identified contain a tyrosine residue, a serine residue, or a threonine residue. Eight-hundred-twenty bisubstrate inhibitors of protein kinases can be generated from these 82 peptides, the ten nucleotides and nucleotide analogs which comprise a triphosphate taught in the specification, a simple 2-carbon tether taught in the specification, and the linking requirements recited in claim 60 (82 peptides x 10 nucleotides or nucleotide analogs x 1 tether = 820 bisubstrate inhibitors of protein kinases). Using a 2-amino-3-(4-amino-phenyl)-propionic acid residue for the tyrosine residue, a 2,3-diamino-propionic acid residue for the serine residue, or a 2,3-diaminobutyric acid residue for the threonine residue (as disclosed at page 8, paragraph 29) in each of the 82 peptides an additional 820 bisubstrate inhibitors of protein kinases (82 peptides x 10 nucleotides or nucleotide analogs x 1 tether = 820 bisubstrate inhibitors of protein kinases) are disclosed. Thus, 820 representative species have been described in the specification.

The attached Declaration under Rule 132 of inventor Philip Cole (Attachment 5) presents data regarding an additional five bisubstrate inhibitors. These were made according to the teachings of the present application. The inhibitors are directed to five protein kinases distinct from those targeted by the inhibitors disclosed in the application. Each was found to be a potent inhibitor of its target enzyme. These data demonstrate that the species disclosed in the application are indeed representative of the claimed genus.

Thus, the specification discloses sufficient identifying characteristics of a representative number of species of the bisubstrate inhibitors of protein kinases and bisubstrate inhibitors of insulin receptor kinase. The specification teaches identifying characteristics of the peptide, of the nucleotide or nucleotide analog moieties, of the linkage between the nucleotide or nucleotide analog moiety and the tether, and of the linkage between the peptide moiety and the tether. One skilled in the art would reasonably conclude that the applicants had possession of the claimed genus of bisubstrate inhibitors of protein kinases and the claimed genus of bisubstrate inhibitors of

insulin receptor kinase when they filed the application. Withdrawal of this rejection is respectfully requested.

## The Rejection of Claims 1-14, 58, 60, 63, 66 and 67 Under 35 U.S.C. § 112, First Paragraph

Claims 1-14, 58, 60, 63, 66 and 67 stand rejected under 35 U.S.C. § 112, first paragraph as allegedly failing to enable the genus of bisubstrate inhibitors of insulin receptor kinase or protein kinases. In particular, the rejection urges that undue experimentation would be required to practice the genus of bisubstrate inhibitors of protein kinase and the genus of bisubstrate inhibitors of insulin receptor kinase. Office Action, Paper No. 04122004, page 12, last paragraph. Applicants respectfully traverse this rejection.

An analysis of whether a claim is enabled by the specification requires a determination of whether the specification contains sufficient information, together with knowledge in the prior art, to enable one skilled in the art to make and use the claimed invention without undue experimentation. "The test of enablement is whether one reasonably skilled in the art could make or use the invention from the disclosures in the patent coupled with information known in the art without undue experimentation." United States v. Telectronics, Inc., 857 F.2d 778, 8 U.S.P.Q.2d 1217 (Fed. Cir. 1988). Factors that may be considered in determining whether experimentation is undue include: (1) the breadth of the claims, (2) the nature of the invention, (3) the state of the prior art, (4) the level of one of ordinary skill, (5) the level of predictability in the art, (6) the amount of direction provided by the inventor, (7) the existence of working examples, and (8) the quantity of experimentation needed to make or use the invention based on the content of the disclosure. In re Wands 858 F.2d 731, 8 U.S.P.Q.2d 1400 (Fed. Cir. 1988). The specification need only describe in detail that which is new or not conventional. Hybritech v. Monoclonal Antibodies, 802 F.2d 1367, 231 U.S.P.Q. 81 (Fed. Cir. 1986).

#### The Breadth of the Claims

The Office Action asserts that undue experimentation would be required to make the genus of bisubstrate inhibitors of insulin receptor kinase and the genus of bisubstrate inhibitors of protein kinases. Specifically, the Office Action asserts that the bisubstrate inhibitors comprise any nucleotide analog having any structure, any peptide substrate, and a tether having an undefined structure. Office Action, page 13, first paragraph. In addition, the Office Action asserts that the moieties can be linked in any manner. *Id*.

The claims have been amended so that the structure of the components and their relationship are better defined. The claims as amended positively recite a nucleotide or nucleotide analog moiety which comprises a triphosphate. The peptide moiety in the amended claims is a substrate for protein kinase or insulin receptor kinase and comprises a tyrosine residue or a 2-amino-3-(4-amino-phenyl)-propionic acid residue (claims 1 and 60). The peptide moiety of claim 60 as amended also can comprise a serine residue, a 2,3-diamino-propionic acid residue, a threonine residue, or a 2,3-diamino-butyric acid residue. The claims as amended recite a tether that links the nucleotide or nucleotide analog moiety to the peptide moiety through a gamma phosphorus on the triphosphate of the nucleotide or nucleotide analog moiety and the tyrosine residue via its phenolic oxygen, the 2-amino-3-(4-amino-phenyl)-propionic acid residue via its aniline nitrogen (claims 1 and 60). The tether of claim 60 as amended links the gamma phosphorus of the triphosphate to the hydroxyl oxygen on the serine residue or threonine residue, or to the 3-amino nitrogen atom on the 2,3-diamino-propionic acid residue or 2,3-diamino-butyric acid residue. Thus, the nucleotide or nucleotide analog moiety and peptide moiety have more defined structures, and the nucleotide or nucleotide analog moiety and peptide moiety are linked to the tether in a more specific arrangement. The breadth of the claims has been limited in these regards.

The claims recite a nucleotide or nucleotide analog moiety comprising a triphosphate. The recited nucleotide or nucleotide analog moiety does not comprise any nucleotide or nucleotide analog moiety having any structure, but rather has a defined structure which comprises a triphosphate. The breadth of the claims is limited in this aspect.

The claims also recite a peptide moiety which is a substrate for an insulin receptor kinase and comprises a tyrosine residue or a 2-amino-3-(2-amino-phenyl)-propionic acid residue (claim 1) or a substrate for a protein kinase and comprises a tyrosine residue, a 2-amino-3-(4-amino-phenyl)-propionic acid residue, a serine residue, a 2,3,-diamino-propionic acid residue, a threonine residue, or a 2,3-diamino-butyric acid residue (claim

60). Thus, the peptide moiety does not comprise any peptide substrate, but rather has a defined composition and function. The breadth of the claims is limited in this regard.

In addition, the claims recite a tether that is greater than 4.9 Å and is linked to the peptide moiety and to the nucleotide or nucleotide analog moiety in a specific arrangement. The tether is limited to a tether that is greater than 4.9 Å. For the bisubstrate inhibitors of insulin receptor kinase, the tether is linked to the tyrosine residue of the peptide moiety via its phenolic oxygen, or is linked to the 2-amino-3-(4-amino-phenyl)-propionic acid residue via its aniline nitrogen. The tether of the bisubstrate inhibitors of protein kinases is linked to tyrosine residue via its phenolic oxygen, the 2-amino-3-(4-amino-phenyl)-propionic acid residue via its aniline nitrogen, the serine or threonine residues via their hydroxyl oxygen, or the 2,3-diamino-propionic acid residue or 2,3-diamino-butyric acid residue via their 3-amino nitrogen. The tether also is linked to the nucleotide or nucleotide analog moiety via a gamma phosphate of the triphosphate. Thus, the tether is not linked to the nucleotide or nucleotide analog moiety and peptide moiety in any manner, but rather is linked in a prescribed manner. The breadth of the claims is limited in this regard.

#### The State of the Prior Art

The rejection asserts that the state of the prior art regarding bisubstrate inhibitors of protein kinases and insulin receptor kinase was not advanced. Office Action, page 14, lines 10-12. The reason stated is that the nucleotide analog and peptide can have any structure. In addition, the nucleotide or nucleotide analog and peptide can be linked to a tether in any arrangement.

As a preliminary matter, the claims have been amended, as detailed above, so that the nucleotide analog and peptide moieties have additional structural requirements. In addition, the claims have been amended so that the nucleotide or nucleotide analog, peptide and tether are linked in a prescribed manner. Thus it is no longer accurate to assert that any structure or arrangement is encompassed.

The state of the prior art was advanced at the time applicants filed their patent application. Nucleotides and nucleotide analogs which comprise a triphosphate were well known in the prior art. Applicants teach nucleotide and nucleotide analogs comprising a triphosphate. Page 7, paragraph 27: "Suitable moieties include ATP, ATPγ-

S, GTP, CTP, TTP, UTP, GTPγ-S, CTPγ-S, TTPγ-S [and] UTPγ-S." Each of the nucleotides or nucleotide analogs contains a triphosphate. In addition, other nucleotides and nucleotide analogs containing a triphosphate were well known in the art and were commercially available to the skilled worker. Examples of such nucleotides and nucleotide analogs include dATP, dCTP, dGTP, dTTP, ITP, dITP, 2',3'-dideoxy-ATP (ddATP), 2',3'-dideoxy-CTP (ddCTP), 2',3'-dideoxy-GTP (ddGTP), 2',3'-dideoxy-TTP (ddTTP), 8-bromo-ATP, 5-bromo-dUTP, 5-iodo-CTP, 5-iodo-dCTP, 5-iodo-UTP, 8-azido-ATP, 5-(3-aminoallyl)-2'-dUTP, 5-(3-aminoallyl)-ATP, and ribavirin-5'-triphosphate. See Attachment 2.

A host of both natural and non-natural peptide substrates were known in the art. As discussed above, applicant's identified in the prior response (dated March 17, 2004) nineteen peptides that were known in the art to be natural peptide substrates of insulin receptor kinase or were known to function as substrates of the insulin receptor kinase. See Attachment 3 for a list of such peptides. Eighty-two additional peptides also were identified in the previous amendment that were known in the art to be substrates of protein kinases. See Attachment 4. Thus, a skilled worker would only need to select from the known nucleotides and nucleotide analogs comprising a triphosphate, and to select from the known natural and non-natural peptide substrates, and to link the two moieties via a tether using the linking requirements taught in the specification and recited in claims 1 and 60. The state of the prior art was advanced at the time of filing with regard to the components for making the inhibitors of the invention.

#### The Skill in the Art

The rejection asserts that the level of skill in the art was insufficient to enable the bisubstrate inhibitors of insulin receptor kinase or protein kinases. The reason stated is that the bisubstrate inhibitors are unlimited with respect to the peptide moiety, nucleotide analog moiety, and tether. Office Action, paragraph bridging pages 14-15.

As a preliminary matter, the claims have been extensively amended so that the structures claimed are not unlimited as asserted.

The level of one of ordinary skill was high at the time applicants filed their patent application. The skilled worker in the field was a protein chemist. Such persons typically have a Ph.D. degree with several years of post-doctoral training. Such persons

would have knowledge of natural and non-natural peptide substrates, and of nucleotides or nucleotide analogs which comprise a triphosphate, as described in the prior art. In addition, the skilled worker could easily link a peptide moiety and a nucleotide or nucleotide analog moiety which comprises a triphosphate in accordance with the linking requirements taught in the specification and recited in claims 1 and 60. See Example 2, page 12. The reactions disclosed are standard coupling reactions run under standard coupling conditions, and are characterized as such at paragraph 41, lines 4-5.

#### Level of Predictability

The rejection asserts that the level of predictability is low because the specification fails to provide any guidance as to the way in which the nucleotide or nucleotide analog moiety, the peptide moiety, and the tether are physically linked.

As a preliminary matter the claims have been extensively amended so that the way in which the component moieties are linked is specified.

Because the prior art was rich and the skill level in the art was high, the level of predictability would also have been high. The specification teaches and claims 1 and 60 recite specific linking requirements for linking the nucleotide or nucleotide analog moiety and the peptide moiety. No reasons have been put forward why these components could not have been predictably joined. No reasons have been put forward why such joined components should not function in the intended manner. In fact, the attached Declaration of Dr. Philip Cole (Attachment 5) presents additional examples where the assembled components of bisubstrate inhibitors do function in the intended manner.

#### The Amount of Guidance

The rejection asserts that the specification fails to provide guidance for the composition and length of the tether. However, the specification teaches that the tether comprises carbon, hydrogen, or oxygen atoms. Page 8, paragraph 29. The length of the tether recited in the claims is greater than or equal to 4.9 Å measured from the gamma phosphorus of the nucleotide or nucleotide analog moiety and a proton donor formed by the phenolic oxygen, aniline nitrogen, hydroxyl oxygen, or 3-amino nitrogen. Guidance in how to calculate the length of the tether is taught at page 4, paragraph 17, lines 4-6. Thus the specification provides guidance for the composition of the tether and for the length of the tether.

#### Quality of Experimentation Needed

The rejection asserts that the amount of experimentation required to practice the invention would be undue because the nucleotide or nucleotide analog moiety and the peptide moiety can have any structure and there is no limitation to where the moieties are linked to the tether. However, as explained above, the claims as amended are not unlimited. The claims have been amended to recite a nucleotide or nucleotide analog moiety comprising a triphosphate, a peptide moiety comprising a tyrosine residue or a 2-amino-3-(4-amino-phenyl)-propionic acid residue (claims 1 and 60), or a serine residue, a 2,3-diamino-propionic acid residue, a threonine residue, or a 2,3,diamino-butyric acid residue. The claims also have been amended to recite a specific arrangement for the linkage of the nucleotide or nucleotide analog moiety and the tether, and the peptide moiety and the tether.

Given the breadth of the claims, guidance of the disclosure, the level of skill in the art, the level of predictability, and the state of the art, one of ordinary skill in the art could have practiced the invention without undue experimentation. All component parts of the claimed bisubstrate inhibitors were known. One of skill would merely need to assemble the parts using the linking requirements recited in claims 1 and 60. Such assembly would be routine and not require undue experimentation.

#### The Rejection of Claims 60, 67, 69-70, and 74 Under 35 U.S.C. § 102(b)

Claims 60, 67, 69-70, and 74 stand rejected under 35 U.S.C. § 102(b) as anticipated by Ricouart *et al.*, *J. Med. Chem.* 34:73-78, 1991. Applicants respectfully traverse this rejection.

To anticipate a claim a reference must teach each and every limitation of the claim. "A claim is anticipated only if each and every element as set forth in the claim is found, either expressly or inherently described in a single prior art reference." *Verdegaal Bros. v. Union Oil Co. of California*, 814 F.2d 628, 2 U.S.P.Q.2d 1051 (Fed. Cir. 1987).

Independent claim 60 and dependent claims 67, 69-70, and 74 are directed to a bisubstrate inhibitor of a protein kinase. The inhibitor comprises (1) a nucleotide or nucleotide analog moiety which comprises a triphosphate and (2) a peptide moiety. A

tether links the moieties. The tether is linked to the nucleotide or nucleotide analog moiety via the gamma phosphate of the triphosphate.

Ricouart is cited, *inter alia*, as teaching bisubstrate inhibitors that have alkyl groups substituted for phosphate groups on nucleotide analogs. "Ricouart et al. teach bisubstrate inhibitors of PKC comprising various ATP mimics having alkyl groups in place of the nucleotide phosphates . . . ." Office Action at page 17, paragraph 11.

Ricouart teaches inhibitors of protein kinase A (PKA) and protein kinase C (PKC). Ricouart, page 74, Table II. Ricouart's inhibitors have an isoquinoline-5-sulfonamide and a Ser-Arg<sub>6</sub> peptide bound together by a linker (-NH(CH<sub>2</sub>)<sub>2</sub>. NH(CH<sub>2</sub>)<sub>2</sub>CO-). Abstract. An exemplary inhibitor taught by Ricouart is shown below.

The Ricouart inhibitors do not contain a nucleotide or nucleotide analog which comprises a triphosphate. Because the Ricouart inhibitors do not contain a triphosphate, the inhibitors also do not have a tether that is linked to the nucleotide or nucleotide analog moiety via a gamma phosphate group.

Applicants' independent claim 60 and dependent claims 67, 69-70, and 74, positively recite a nucleotide or nucleotide analog moiety "comprising a triphosphate." In addition, claim 60, and claims 67, 69-70, and 74, positively recite that a tether is linked to the nucleotide or nucleotide analog moiety via "the gamma phosphate of the triphosphate." Ricouart does not teach these two claim limitations. Thus, Ricouart cannot anticipate claims 60, 67, 69-70, and 74. Withdrawal of this rejection is respectfully requested.

Respectfully submitted,

Dated: September 29, 2004

By:

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#### Attachment 1

#### 2-amino-3-(4-amino-phenyl)-propionic acid

$$H_2N$$
 OH  $NH_2$ 

#### 2,3-diamino-propionic acid

#### 2,3-diamino-butyric acid

$$H_2N$$
 OH  $NH_2$ 

#### tyrosine

$$H_2N$$
 OH OH

#### serine

$$H_2N$$
 OH OH

#### threonine

# BIOCHEMICALS AND REAGENTS FOR LIFE SCIENCE RESEARCH

1998

NEW Products

ALPHABETICAL

BIOACTIVE PEPTIDES

IMMUNO-CHEMICALS

MOLECULA Biology

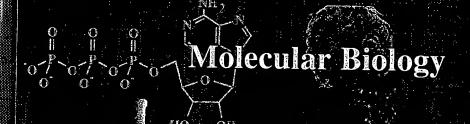
> TISSUE CULTURE

OTHER PRODUC GROUPS

EQUIPMEN Books and

DIAGNOSTIC KITS AND REAGENTS

PRODUC



SIGNAL TRANSDUCTION

cell culture



IMMUNOCHEMICALS



SIGMA

## ALPHABETICAL LIST OF COMPOUNDS

ontinued)	AMINO ACIDS Buch	US \$   PRODUCT   NUMBER
Aldino 27	AMINO ACIDS, Protected (continued)	AMINOACYL-IRNA SYNTHETASE
	VALINE NATROCA Valina A	I IEL I IPASA SUB-clase C 1 1)
5 an Page 197	N-t-BOC-0-Valine Page 199 N-t-BOC-0-Valine Page 199	J VIJI Detinition: One unit will and
an ruge	N-I-DUU-I-Valine Page 100	1.0 picomole (10 <sup>-12</sup> mole) of labeled amino acid to
į.	N-UBZ-DL-Valine Page 252	tRNA in 10 min at pH 7.6 at 37°C (amino acid used:
age 199	N-CBZ-t-Valine Page 252	Protein determined by Biuret method.
	N-CBZ-L-Valine p-Nitrophenyl Ester 0	[9028-02-8]
	I U U U U U U U U U U U U U U U U U U U	
ge 253	N-FMOC-t-Valine Page 491	A mixture of a mix
	N-o-Nitrophenyisulfenyl-Lvaline Page 805 L-Valinamide Page 1122	activating enzymes in 50%
er Page 489	L-Valine Benzyl Ester Page 1122	Blycerol solution containing
5	L-Valine t-Birtyl Ester Page 1122	I IV MM ITIS HCI DH 7.2 10 mM M - OL DO
i i	L-Valine Ethyl Ester Page 1122	2-mercaptoethanol and 10 mM KCl.
	DL-Valine Methyl F ster Page 1100	Accusity: 2,000-6,000 units per mg protein
ı	L-Valine Methyl Ester Page 1123	7 3316 Crude: From Rovine Liver 10 000
1	MISCELLANEOUS	
	Nt-BOC-Amino Acid Resin Esters Page 192	
		glycerol solution containing 10 mM Tris HCl, pH 7.6,
	INT-DUCT-Q-AMINODUITORIC ACID Date 100	KCI.
	ITTOUGTV-MITHIODHING ASIA D 100	Activity: 2,000-7,000 units per mg protein.
	ITTOUG-7-AMINONENTANCIC Acid D 100	7 3040 Crude: From F coll
		A mixture of amino acid 10,000 units 39.23
	Acid Page 103	activating enzymes in FOOV
195	N-t-BOC-(2S, 3R)-3-Aming-2-hydroxy 4 phonythal	Blycerol solution containing
1		1 IO IIIN INS HOLDER 7.5
nimide 📲	N-t-BOC-bu-2-(t-Butyl)glycine Page 194	10 mM MgCl <sub>2</sub> , 30 mM 2-mercaptoethanol and 10 mM KCl.
. 3	11770007-HUHDSPING Page 104	Activity: 10 000 15 000
	11-004-7-AMMO-0-hithric Acid D	Activity: 10,000-15,000 units per mg protein.
No.		8-AMINOADENOSINE 3':5'-CYCLIC 2 mg 50.20 A 4637 MONOPHOSPHATE 5 mg 92.95
4	N-CBZ42R,3R)-3-Amino-2-hydroxy-4-phenylbutyric 단본Acid Page 248	Free Acid 5 mg 82.85
ter Page 199	N-CBZ-D-3-(2-Nanhthyl)alanina R	[30685-40-6] C <sub>10</sub> H <sub>13</sub> N <sub>6</sub> O <sub>6</sub> P FW 344.2
ž.		
er Page 199	No-Nitrophenylsulfenyl-y-aminobutyric Acid Page 804	β-AMINOADIPIC ACID A 1786 (3-Aminohexanedioic acid)
	E)	- 140967-78-01 C.H. NO. DV 151 0
	AMINO ACID SOLUTIONS	R: 36/37/38 S: 26-36
253	See: Tissue Culture Media and Description	AREAN A PURIS
	See: Tissue Culture Media and Reagents Page 1752	
enyl	AJ: AMINO AOIT	Glutamine synthetase inhibitor in 250 mg 37.95
	A.S. AMINO ACID STANDARD SOLUTIONS See under Protein Applying Protein	VIVO. dilling acid antagonist
ge 490	See under: Protein Analysis Reagents Page 2115	1 [/020-28-2] C <sub>c</sub> H <sub>2</sub> ,NO 6 210.30
5	A CONTRACTOR OF THE CONTRACTOR	
osine Page 806	9-AMINOACRIDINE	A 0637 Minimum 999 100 mg 6.80
	(Aminacrine)	14 003/ Minimum 99%
A STATE OF	A7295 Free Base	The duces kalliale toxicity, alithamina
1	Produces a hazy colution in ut 3 g 23.75	5 [542-32-5] C.H. NO. THE 5 g 98.90
V.	190-45-91 Co.H. N. EW 104-9. 25 g 76.25	(5-2-02-0) C6H11NO4 FW 161.2 10 g 179.10
7 W		A 7275 (L-2-Aminophyspacific said) 100 mg 14.40
	26-36/37/39-22 S: 45-	
A STATE OF THE STA	A1135 FU.	antagonist increases into 1 g 79.80
		free Ca <sup>2+</sup> : competitive inhibit
	53217-22-8] C <sub>13</sub> H <sub>10</sub> N <sub>2</sub> • HCl FW 230.7 23.7 23.80 25 g 29.80	glutamine synthetase and auglutamide sets
	FW 230.7 LI3H10N2 • HCI	
2	R: 23/24/25-36/27/20 40 0 45 0	[1118-90-7] C <sub>6</sub> H <sub>11</sub> NO <sub>4</sub> FW 161.2
	\$3,53,738-40 S: 45-26-36/37/39-22	5-(3-AMINOALLYLL-2'-DEOVY
4		WOULD ORIDINE 5-TRIPHOSPHATE
	744MINOACTINOMYCIN D 1 mg 66.35 mg 245.65 mg 245.65	(AA-dUTP) 3 III 266./0
	Transfer of the contract of th	ovalum San
	18:45-46-61-26/27/28 St 45-26-26 72 72	Approx. 90%
	Ri4546-61-26/27/28 S: 45-26-36/37/39	[ $109921-28-0$ ] $C_{12}H_{20}N_3O_{14}P_3$ FW 523.2 (for free acid)
	AMINOACY	
4	ACVISCA LO	5-(3-AMINOALLYL)URIDINE 5'-TRI- 1 mg 80.10
(F	Acylase I Page 53	-OC (AA LITE) 5 mg 266 70
		(AA-UTP) 5 mg 266.70 Sodium Salt 10 mg 444.40
<b>3</b>	CIMINOACVI	Approx. 80%
	CHANGE TO THE PART OF THE PART	- TP1 VA. UU/0
	Prolidase Page 934	[7522]-88-4] C <sub>12</sub> H <sub>20</sub> N <sub>3</sub> O <sub>15</sub> P <sub>3</sub> FW 539.2 (for free acid)

### **ALPHABETICAL**

3'-A7IDO-3'-DEOXYTHYMIDING 2 140

PRODUCT			us s	PRODUCT NUMBER			us \$
NUMBER 6.	-AZAURIDINE	250 mg	9.30		8-AZIDOADENOSINE 5'-TRI-	1 mg	26.05
A.1882	(6-Azauracil riboside)	1 g	21.95		PHOSPHATE Sodium Salt	5 mg 10 mg	85.65 142.55
2-8°C	See also: <b>3-Deazauridine</b> Page 349	5 g	67.00	1 <b>♦</b> .	Minimum 75%	Shipped in	
	[54-25-1] C <sub>8</sub> H <sub>11</sub> N <sub>3</sub> O <sub>6</sub> FW 245.2		,	*	Light tan powder. Useful in photoaffinity labeling.		
	R: 20/21/22-40 S: 22-36		)	· ~	Ref.: Czarnecki, J., et al., Meth. Er	nzymol., 56	, 642
	B-AZAXANTHINE (8-Aza-2,6-dihydroxypurine)	1 g	9.80	1	(1979). [53696-59-6] C <sub>10</sub> H <sub>15</sub> N <sub>8</sub> O <sub>13</sub> P <sub>3</sub> FW		
A 2132	Minimum 85%		)	ı	acid)		
	[1468-26-4] C <sub>4</sub> H <sub>3</sub> N <sub>5</sub> O <sub>2</sub> FW 153.1		1		4-AZIDOBENZOIC ACID	50 mg	29.40
A	AZELAIC ACID (Nonanedioic acid)		1	A 9048	N-HYDROXYSUCCINIMIDE ESTER	100 mg 500 mg	49.00 213.40
	[123-99-9] C <sub>9</sub> H <sub>16</sub> O <sub>4</sub> FW 188.2		,	1	(N-Hydroxysuccinimidyl	<b></b>	•••
	R: 36/37/38 S: 26-36		!	ĺ	4-azidobenzoate) Minimum 95%		
A 2282	Approx. 98%	1 g 10 g	5.45 9.55	1	A photoactivatable, heterobifunction	onal cross-lir	nking
(RY)		25 g	19.10		reagent Ref.: Galardy, R.E., et al., J. Biol. (	Chem., 249	3. 3518
		100 g	45.00	1	(1974).		,
A 2257 RT		5 g 25 g	5.40 8.10	1	[53053-08-0] C <sub>11</sub> H <sub>8</sub> N <sub>4</sub> O <sub>4</sub> FW 26		
(R.I.)		100 g	11.40		8-AZIDO-CYCLIC ADENOSINE DIPHOSPHATE-RIBOSE	Nucle 100 µg	eotide 376.20
		500 g 1 kg	20.05 36.15	C-14-1-1	Minimum 95% (HPLC)	,0	
	410 40ID CADDOYV-14C			1	Lyophilized powder containing sodium chloride		
,	AZELAIC ACID-CARBOXY-14C  See: Radiochemicals Section Page 2	2123		1	Photoaffinity labelled; analog of cy	yclic ADP-rib	ose
	AZELAOYL CHLORIDE		11.60	1	Ref.: Walseth, T.F. and Lee, H.C., Acta, 1178, 235 (1993).		орпуь.
A 7436	(Nonanedioyl dichloride)			1	[150424-94-5] C <sub>15</sub> H <sub>20</sub> N <sub>8</sub> O <sub>13</sub> P <sub>2</sub> F	FW 582.3	
<u>-o·c</u>	Approx. 98% (GC) [123-98-8] C <sub>9</sub> H <sub>14</sub> Cl <sub>2</sub> O <sub>2</sub> FW 225.1	ı			2'-AZIDO-2'-DEOXYCYTIDINE	10 mg	70.15
*	R: 34 S: 26-28-27-36/37/39			A 7783	3 May contain up to 5% inorganic salts.		
	L-AZETIDINE-2-CARBOXYLIC	50 mg	23.25		[51034-68-5] C <sub>9</sub> H <sub>12</sub> N <sub>6</sub> O <sub>4</sub> FW 26	68.2	
A 0760		100 mg 250 mg	38.70 77.40		3'-AZIDO-3'-DEOXYTHYMIDINE	25 mg	24.35
RT.	Crystalline		215.00		9 (AZT; Azidothymidine)	100 mg 250 mg	67.35 148.10
	A four-membered ring analog of L-proline.			1-00	l a	-	
	[2133-34-8] C <sub>4</sub> H <sub>7</sub> NO <sub>2</sub> FW 101.1			_	HN CH	3	
	8-AZIDOADENOSINE	100 mg	21.90	,	رير <sub>ا</sub> م		
	[4372-67-2] C <sub>10</sub> H <sub>12</sub> N <sub>8</sub> O <sub>4</sub> FW 308.3				HOCH <sub>2</sub>		
		5 mg	27.80	, l	<u></u>		
	8-AZIDOADENOSINE 3':5'-CYCLIC MONOPHOSPHATE	5 mg	27.80		N <sub>3</sub>		
-oc	Free Acid				[30516-87-1] C <sub>10</sub> H <sub>13</sub> N <sub>5</sub> O <sub>4</sub>		
	Approx. 95% Reported to be of use in photoaffini	ity labeling			FW 267.2 R: 23/24/25 S: 45-36		
	Ref.: Haley, B.E. and Hoffman, J.F. Sci. USA, 71, 3367 (1974).	., Proc. Nat	d. Acad.		AZIDOTHYMIDINE, ANTIBODY TO		
_	[31966-52-6] C <sub>10</sub> H <sub>11</sub> N <sub>8</sub> O <sub>6</sub> P FW 3	370.2			See: Immunochemicals Page 136	61	
	8-AZIDOADENOSINE 5'-DI-	1 mg	22.10		3'-AZIDO-3'-DEOXYTHYMIDINE-2		
A 6657	PHOSPHATE	5 mg	80.15 140.10	5	See: Radiochemicals Section Pag		19.05
-0°C	Approx. 95%	10 mg	140.10	A 0679	3'-AZIDO-3'-DEOXYTHYMIDINE 79 β-D-GLUCURONIDE	5 mg 25 mg	87,55
	Off-white powder. Reported to be useful in photoaffini	heling	-	RT RT	(AZT glucuronide)	100 mg	
	Reported to be useful in photoaffini Ref.: Czarnecki, J., et al., Meth. Er				Sodium Salt Minimum 97% (HPLC)		
	(1979). [102185-14-8] C <sub>10</sub> H <sub>14</sub> N <sub>8</sub> O <sub>10</sub> P <sub>2</sub> F	-			1133525-01-61 CueHooNeOuoNa	FW 465.4	- 22
	[102185-14-8] C <sub>10</sub> H <sub>14</sub> N <sub>8</sub> U <sub>10</sub> P <sub>2</sub> F acid)	W 400	Ji no-		R: 23/24/25-36/37/38 S: 45-2		
	8-AZIDOADENOSINE 5'-MONO-	1 mg			3'-AZIDO-3'-DEOXYTHYMIDINE-N See: Radiochemicals Section Page	<b>∥ETHYL- 11</b>	
A 8141	PHOSPHATE	5 mg	88.05	5	3'-AZIDO-3'-DEOXYTHYMIDINE	25 mg	115.25
-o·c	Ammonium Salt Approx. 95%	10 mg	146.55	A 680	06 5'-MONOPHOSPHATE	50 mg	001 6h
	Off-white powder.	S. S. Sim		-o°C	(AZT monophosphate)		
	Reported to be of use in photoaffin Ref.: Haley, B.E. and Hoffman, J.F.	ity labeling Proc. Na	i. atl Acad		Sodium Salt Approx. 98%		- 1
	Sci. USA, <b>71.</b> 3367 (1974).				[29706-85-2] C <sub>10</sub> H <sub>14</sub> N <sub>5</sub> O <sub>7</sub> P FW	N 347.2 (for	r free
	[ $102185 \cdot 18 \cdot 2$ ] $C_{10}H_{13}N_8O_7P$ FW acid)	/ 388.2 (tor	r free		acid) R: 20/21/22 S: 36		لب
	aciu						

	3'-AZIDO-3'-DEOXYTHYMIDINE-2-	.14C
	5'-MONOPHOSPHATE See: Radiochemicals Section Page	2123
	2'-AZIDO-2'-DEOXYURIDINE	
A 1021		25 mg
-0°C	FW 269.2	
	R: 36/37/38 S: 26-36	_
	3'-AZIDO-2',3'-DIDEOXYURIDINE	10 mg
A 4810		50 mg
-0°C	Inhibitor of HIV replication Ref.: 1. Lin, T-S. and Mancini, W.R	I Mod (
	<b>26.</b> 544 (1983).	
	2. Zhu, Z., et al., Mol. Pharmacol.,	38, 929 (1
	[84472-85-5] C <sub>9</sub> H <sub>11</sub> N <sub>5</sub> O <sub>4</sub> FW 25	3.2
	AZIDOFLUORESCEIN DIACETATE	25 mg
A 4511 2-8°C	(Azido-FDA) Approx. 95% (HPLC)	
	A photolabelling reagent in membra	ane viscosil
	studies.	
	Ref.: Rotman, A. and Heldman, J., 5995 (1981).	Biochem.,
	[77162-07-3] C <sub>24</sub> H <sub>15</sub> N <sub>3</sub> O <sub>7</sub> FW 45	7 4
	I-(5-AZIDO-2-NITROBEN-	
A 3282	ZOYLOXY)SUCCINIMIDE	50 mg 100 mg
2-8°C	Approx. 95%	_
	Photoactive, heterobifunctional cros Ref.: Lewis, R.V., et al., Biochemist	ss-linking re
	(1977).	ry, 10, 56
	[60117-35-3] C <sub>11</sub> H <sub>7</sub> N <sub>5</sub> O <sub>6</sub> FW 305	.2
6	-(4-AZIDO-2-NITROPHENYL-	50 mg
A 3407 -ॐc	AMINO)HEXANOIC ACID	100 mg
120	N-HYDROXYSUCCINIMIDE ESTER	
	(N-Succinimidyl 6-[4-azido-2-nitroani	linolhevano
	MINIMUM 95%	
	Photoactive, heterobifunctional cros Ref.: Ballmer-Hofe, K., et al., Anal. E	s-linking re
	240 (1983).	
	[64309-05-3] C <sub>16</sub> H <sub>18</sub> N <sub>6</sub> O <sub>6</sub> FW 390	).4
N	-(4-AZIDO-2-NITROPHENYL)-	500 µg
A 1935	N'-(3-BIOTINYLAMINO-	1 mg
	PROPYL)-N'-METHYL- 1,3-PROPANEDIAMINE	2 mg 2
	(Photobiotin)	
	Acetate Salt	
	Minimum 98%	
	Photoactive reagent for covalent mo- biotin.	dification w
	0	
: .	Ĭ	
	ни н	
•	H N	
	s H	
	N N N C	H <sub>3</sub>
	\ \ \ \ \ \	3
	N <sub>3</sub> NO <sub>2</sub>	
	See also: Molecular Biology Products [96087-37-5] C₂₃H₃₅N₀O₄S • C₂H₄O₂	Page 1644
. (2)	S,3R,4E)-2-AZIDO-	
A 0456	A COTE - Z-AZIDO-	1 mg

A 0456 4-OCTADECENE-1,3-DIOL  $1 \; \mathrm{mg}$ (b-Sphingosine azide) [103348-49-8] C<sub>18</sub>H<sub>35</sub>N<sub>3</sub>O<sub>2</sub> FW 325.5

12-AZIDOOLEIC ACID A 8265
Photosensitive chemical for studies of phospholipid-protein 100 mg 500 mg studies of phospholipid-protein interactions in biological membranes **Ref.**: Chakrabarti, P. and Khorana, H.G., Biochemistry, **14**, 5021 (1975). [57818-47-0] C<sub>18</sub>H<sub>33</sub>N<sub>3</sub>O<sub>2</sub> FW 323.5

PRODUCT NUMBER			US \$	PRODUCT NUMBER			USS
	BROMOACETOPHENONE (Phenacyl bromide) Recrystallized, white to light yellow crystals. Suitable for the derivatization and subs	1 g 5 g 10 g	4.95 15.65 25.95 HPLC		BROMOADENOSINE 5'-MONO- PHOSPHATE Free Acid Approx. 98% [23567-966] C <sub>10</sub> H <sub>13</sub> BrN <sub>5</sub> O <sub>7</sub> P FW	5 mg 25 mg 426.1	14.65 46.40
	analysis of fatty acids Use-tested. Ref.: 1. Wood, R. and Lee, T., J. Chron 237 (1983). 2. Mentasti, E., et al., J. Chromatogr., (1985).   70-11-1  CeH,BrO FW 199.0 R: 34 S: 26-27-36/37/39	matogr.,	254,	B 3756	BROMOADENOSINE 5'-TRIPHOS-	5 mg 25 mg / 586.1 (fo	12.30 35.35
	ROMOACETYL-CELLULOSE See under: Affinity Chromatography Me				6α-BROMOANDROSTERONE (5α-Androstan-16α-bromo-3α-ol- 17-one)	1 mg 5 mg	33.65 122.90 220.95
	ROMOACETYL CHLORIDE Approx. 95% (NMR) May contain chloroacetyl chloride and bromoacetyl bromide. [22118-098] C <sub>2</sub> H <sub>3</sub> BrClO FW 157.4 R: 34-14 S: 26-27-36/37/39-3/7/9	50 g 100 g	35.55 59.25	1 B 9392	3α-ol-17-one)	5 mg	117.60
	-BROMOADAMANTANE [768-90-1] C <sub>10</sub> H <sub>15</sub> Br FW 215.1	25 g	18.90	B 0755	[115115-49-6] C <sub>19</sub> H <sub>29</sub> BrO <sub>2</sub> FW 36 •BROMOANILINE (2-Bromoaniline) Yellow to brown solid 1615-36-11 C <sub>8</sub> H <sub>4</sub> BrN FW 172.0		27.85
		00 mg 00 mg 1 g	14.45 45.00 74.10		R: 20/21/22-36/37/38 S: 45-26-3 <b>BROMOANILINE</b> (3-Bromoaniline) d = 1.58 g/ml	36/37/39- 25 ml 100 ml	56.20
8 B 6272 		1 g	12.60 31.80 125.60	p.	[59]: $1.9-5$ ] $C_6H_6BrN$ FW 172.0 R: $20/21/22-36/37/38$ S: $45-26-3$ <b>BROMOANILINE</b> (4-Bromoaniline) [ $106-40-1$ ]. $C_6H_6BrN$ FW 172.0 R: $20/21/22-36/37/38$ S: $45-26-3$		
8	-BROMOADENOSINE 3':5'-CYCLIC N PHATE (8-Br-cAMP) Membrane permeable cAMP analog; re hydrolysis by phosphodiesterases			B 2395 BT  ◆ B 2752		10 g 25 g 50 g 100 g	17.80 35.50 55.35 90.60 24.25
B 5386 	[23583-48-4] C <sub>10</sub> H <sub>11</sub> BrN <sub>5</sub> O <sub>6</sub> P 2 FW 408.1 10	5 mg 25 mg 00 mg 50 mg	13.35 44.50 123.60 296.65	3	Tan powder, may produce turbid solutions.  -BROMOBENZALDEHYDE  [31/32-99-8] C.HsBrO FW 185.0	100 g	62.65
B 7880 -oc	Sodium Salt Approx. 98% 2 [7693946-3] 10	5 mg 25 mg 00 mg	13.60 42.30 117.45 276.70	4	R: 36/37/38 S: 26-36  -BROMOBENZALDEHYDE [1122-91-4] C <sub>7</sub> H <sub>5</sub> BrO FW 185.0 R: 22-36/37/38 S: 26-36	25 g	38.20
	See also: Nf,2'-0-Dibutyryladenosine 3':5'-Cyclic Monophosphate Page 375 Nf-Monobutyryladenosine 3':5'-Cyclic Page 763 2'-0-Monobutyryladenosine 3':5'-Cyclic Page 763		B B 8006	<b>ROMOBENZENE</b> d = 1.49 g/ml See olso: Environmental Standards Page 2017 [108-86-1] $C_6H_6Br$ FW 157.0 R: 10-38-51/53 S: 61	100 ml 500 ml 1 liter	7.25 19.70 37.65	
	Monophosphate <i>Page 763</i> 2'-0-Monobutyryl-8-bromoadenosine 3 Monophosphate <i>Page 763</i> N <sup>2</sup> -Monobutyrylguanosine 3':5'-Cyclic l	3′:5′-Cycl		B 2632	ROMOBENZENE-d <sub>5</sub> 99+ atom % D [4165-57-5] C <sub>6</sub> D <sub>5</sub> Br FW 162.0 R: 10-38 S: 24	5 g	34.15
	Page 763 2'-O-Monobutyrylguanosine 3':5'-Cyclic Monophosphate Page 763BROMOADENOSINE 5'-DIPHOS-	c 5 mg	53.35	P B 1134 RT ◆	BROMOBENZENESULFONYL CHLORIDE Crystalline (98-58-8) C <sub>6</sub> H <sub>4</sub> BrClO <sub>2</sub> S FW 255.5 R: 34 S: 26-27-36/37/39	25 g	28.40
B 3881 	PHATE Sodium Salt Approx. 95% [102185-47-7] C <sub>10</sub> H <sub>14</sub> BrN <sub>5</sub> O <sub>10</sub> P <sub>2</sub> FW free acid) R: 23/24/25-36/37/38 S: 45-26-36		192.50 (for	B 6381	-BROMOBENZOIC ACID	25 g	13.55
206	◆ Shipping informa		age 5.	He	ow to use catalog - page 2.		

NUMBER		
B 2884	BROMOBENZOIC ACID (3-Bromobenzoic Acid) Crystalline [585-76-2] C <sub>7</sub> H <sub>5</sub> BrO <sub>2</sub> FW 201.0	5 g 1
	R: 36/37/38 S: 26-36	
р- В 2634 Гос	BROMOBENZOIC ACID (4-Bromobenzoic Acid) Crystalline [586-76-5] C₁H₂BrO₂ FW 201.0 R: 22-36/37/38 S: 26-36	10 g 1
	{2-BROMOBENZYL}-N-2-(CHLOROE	THYI 1-
N·	ETHYLAMINE See: N-(2-Chloroethyl)-N-ethyl-2-bromol Page 269	
BI	ROMOBIMANE 2	5 mg 4
B 4380 ਕਾਫ	MInimum 97% Fluorescent probe for thiols Ref.: 1. Kosower, N.S., et al., Proc. N USA, 76, 3382 (1979). 2. Danielsohn, P. and Nolte, A., Histoc (1987). [71418-44-5] C <sub>10</sub> H <sub>11</sub> N <sub>2</sub> O <sub>2</sub> Br FW 271	hem., <b>86,</b>
A.	BROMO-2-BUTENOIC ACID METHYL	. 5 ml 3
B 3152 2-8°C ◆	ESTER Approx. 97% (GC) d = 1.51 g/ml Stabilized with silver wool. [111771-1] C <sub>5</sub> H <sub>3</sub> O <sub>2</sub> Br FW 179.0 R: 34-42/43 S: 26-27-36/37/39	
3	-BROMO-3-BUTEN-1-OL	1 g 1
41,088-6 2-8°C		10 g 1(
	-(4-BROMOBUTYL)PHTHALIMIDE	10 g 5
B 3502		
2	-BROMOBUTYRIC ACID	100 ml :
B 0136 2-8℃ ◆	d = 1.56 g/ml [80-58-0] C <sub>4</sub> H <sub>7</sub> BrO <sub>2</sub> FW 167.0 R: 23/24/25-34 S: 26-45-27-36/37	
4	-BROMOBUTYRIC ACID	5 g :
B 3627 □==□ ◆	Approx. 98% Yellow to brown semi-solid. [2623-87-2] C <sub>4</sub> H <sub>7</sub> BrO <sub>2</sub> FW 167.0 R: 34 S: 26-27-36/37/39	10 g ; 50 g (
	-BROMOBUTYRIC ACID ETHYL	25 ml
B 3652		
	I-BROMO-CALCIUM IONOPHORE	1 mg
B 7272 2-8°C	A23187 Free Acid Ref.: Debone, M., et al., Biochemistry (1981). [76455-82-8] C <sub>29</sub> H <sub>36</sub> BrN <sub>3</sub> O <sub>6</sub> FW 60 R: 20/21/22-36/37/38 S: 26-36/3	5 mg 3 y, <b>20</b> , 686 2.5
B 6884	[IR]-endo)-(+)-3-BROMOCAMPHOR (3-Bromo-d-camphor) [10293-06-8] C <sub>10</sub> H <sub>15</sub> BrO FW 231.1 R: 36/37/38 S: 26-36	10 g 50 g 100 g
B 1396	([1S]-endo)-(-)-3-BROMOCAMPHOR	

To place an order call 800-325

/ 364.6

	PRODUCT NUMBER				PRODUCT		
	B 0639	ROMOCHLOROMETH d = 1.99 g/mi	ANE	100 g 12.55	NUMBER		
	<b>厨</b> .	See also: Environment	tal Standards Page	2015 and	B 2381		E 25 mg 92
		[74-97-5] CH <sub>2</sub> BrCl   R: 20-41-37/38 S: 2	744 * * * * * * * * * * * * * * * * * *		1	Approx 98%	
	1-	BROMO-3-CHI ODOD				[88188-03-8] C <sub>9</sub> H <sub>13</sub> BrN <sub>3</sub> O <sub>7</sub> P acid)	
7.	2.1	See: Molecular Biology BROMO-1-CHLOROPE	Reagents Page 1.	599	B 1886	5-BROMO-2'-DEOXYCYTIDINE 5'-TRIPHOSPHATE	1 mg 19
_		See: Environmental State Page 2018	<b>IOPANE</b> ndards <i>Page 2013</i>	5, and	<u>⊸</u>	Sodium Salt	5 mg 64.
	2-B	ROMO-2-CHLORO-			<b>A</b>	[30419-11-5] C <sub>9</sub> H <sub>15</sub> BrN <sub>3</sub> O <sub>13</sub> P <sub>1</sub> acid)	FW 546.1 (for free
<u>-</u>		,1,1-TRIFLUOROETH, falothane)		g 16.15 g 16.85		R: 36/37/38 S: 26-36	
•	· N	linimum qqv	250	g 54.25	5-	BROMO-2'-DEOXYURIDINE	
	u	halation anesthetic = 1.88 g/ml		1			
184. 135	" St	abilized with a and a	moi.	1		Thymidine analog used as a mul research.	tagen in genetic
		51-67-7] C <sub>2</sub> HBrClF <sub>3</sub> 20-41-40 S: 26-36-2				0	
	38-B	ROMO-5-CHOLESTEN	3			Ĭ,	).
	Sec	: Cholesteryl Bromide	IE Page 202	-1		HN T	,
3	BRON	OCONDUBITOR				0 N	
)::: 88	. See	6-Bromo-4-cyclohexe	De-1 2 3 trial D			HOCH <sub>2</sub>	
				₹209			
Purch	see.	Bromcresol Green Page	ge 204		15	HO <sup>N</sup>	
16.10	BROM	OCRYPTIME MEONS			R:	9-14-3] C <sub>9</sub> H <sub>11</sub> BrN <sub>2</sub> O <sub>5</sub> FW 307 46-61-20/21/22 S: 45-36/37	.1
	See:	2-Bromo-α-ergocryptin	ie Methanesulfona	. B9	285 Sie	smaUltra	//39
					J Min	Nimum 000/	50 mg 9.50
B 541	8-BRO	MO-CYCLIC ADENOSI OSPHATE RIBOSE	NE 250 μg	41.65	Kes	Sidue after ionia:	250 mg 31.70 1 g 87.80
	(Br-c/	DP-ribose)	500 μg	79.05	СОП	nolete coloriana	1 g 87.80
12 g	90-9	5% /HDI CL	4		mso	NUDIE matter: -0 16/	
		ared enzymatically		- 1	n, <	0.0005% <0.001%	Na: <0.005%
	Phosp	nilized powder contain thate buffer salts.	ning 10-20% soc	lium	Cu; <	<0.0005%	NH4*: <0.05%
	Ref	Pinet of cADP-ribose-inc	luced Ca <sup>2+</sup> release	. 1	re: < K· -∩	0.0005% 0.005%	P: <0.005% Pb: <0.001%
	See als	I. Biochim. Biophys. Actor Cyclic Adenosine Dip 32	ta, 1178, 235 (1	993).	Mg: <	:0.0005%	Zn: <0.0005%
				B 500		mum 99%	
11.1%	R: 36/3	98-26-9] C <sub>15</sub> H <sub>20</sub> BrN <sub>5</sub> 0 37/38 S: 26-36	<sup>13</sup> P₂ FW 620.2	-00			100 mg 10.65
6	-BROMC	-4-CYCLOHEXENE-					250 mg 16.15 500 mg 26.85
1147			1 mg	8.85			<sup>1</sup> g 44.75
	Mixed I	onduritol)		29.25	5-BROM	IO-2' DEOXOGO	5 g 167.95
	Glucosid	aco inhihu	· ·	18.70	See: R	IO-2'-DEOXYURIDINE-2-14C adiochemicals Section Page 21	
	ner: i	Moth E	E. 693 (1007)	- 1	5-BROM	0.3/ Prove	24
	6/87 (1d	1821	ACAG. Sci. USA 7	79.	ANTIB	0-2'-DEOXYURIDINE; MONOC	LONAL
<u> </u>	[42014]	74-4] C <sub>6</sub> H <sub>9</sub> BrO <sub>3</sub> FW 2	100 O	" ].	(Anti-Br	rd(1)	
				_	See: III	munochemicals Page 1280	
	<b>シレロか-</b> おん	-21 CH BAL-	100 mg 23	3.25 B 2506	-BROMO	0-2'-DEOXYURIDINE	
				-o€	Sodium		mg 59.15 mg 97.90
752 <i>[</i> :	<b>KUMOC</b> 1 2240.25	YTOSINE 7] C4H4BrN3O	250 mg 19.	05	-151432	32-71 CH DHA - 25	mg 194.75
Fi Fi	W 190.0	1 C4H4RLN3O	mg 19.	.00	R: 40 S		
10.		ECAN-1-OL					
66 A			1 g 19.	25 B 0631 5.	5'-TPIPH	2'-DEOXYURIDINE 1 IOSPHATE 1	mg 19.75
	~ 1.090	/mi	10 g 130 :	30 ক্রি	Sodium	Sah 5	mg 64 15
R:	36/37/3	6) C <sub>10</sub> H <sub>21</sub> BrO FW 23 8 S: 26-36	7.2	1 3/	Approx.	90% 10	mg 106.85
E 0.		0. 20-30		1	FW 547.0	99.7) C <sub>9</sub> H <sub>14</sub> BrN <sub>2</sub> O <sub>14</sub> P <sub>3</sub> (for free acid)	· <del>-</del>
	022.70.2	DEOXYCYTIDINE C9H12BrN3O4	100 mg 14.6	5	₹: 23/24/	(tor free acid) /25-36/37/38-40 S: 45-26-36	
6 110				~ /			_22
FW	306.1 10 S: 22	O3112D1143O4	500 mg 46.1 1 g 76.2		JMODIC:	HLOROMETHANE	-22

## **ALPHABET**

To place an order call 8

PRODUCT NUMBER			US S	PRODUCT NUMBER		Line	PRODUCT NUMBER
D 9670	3-DEOXY-o-GLYCERO-o-GALAC 2-NONULOSONIC ACID (KDN) Ammonium Salt A natural deaminated sialic acid	50 mg	46.65 184.90	1 :	2'-DEOXYGUANOSINE 5'-MONOPHOSPHO- MORPHOLIDATE 4-Morpholine-N,N'-dicyclohexy salt	100 mg 36.20 1 g 200.55	1-DEOXY-1-MORPHOLINO- D 6149
	identification and quantification residues in poly(oligo)nonuloso Ref.: 1. Nadano, D., et al., J. E 11550 (1986). 2. Kitajima, K., et al., Anal. Bior	nates. liol. Chem., <b>261</b>	١,	2	[102783·39·1] C <sub>14</sub> H <sub>21</sub> N <sub>6</sub> O <sub>7</sub> P • C FW 709.8 S: 7-22 ''-DEOXYGUANOSINE 5'-TRI-	10 mg 24.75	1-DEOXY-1-NITRO-p-MANNITOL D 3651 [14199-83-8] C <sub>6</sub> H <sub>13</sub> NO <sub>7</sub> 配 FW 211.2
	(1992). [112543-66-5] C <sub>9</sub> H <sub>16</sub> O <sub>9</sub> • NH			D 4010 	PHOSPHATE Sodium Salt Approx. 97% [93919-41-6] C <sub>10</sub> H <sub>16</sub> N <sub>5</sub> O <sub>13</sub> P <sub>3</sub>	25 mg 49.20 100 mg 142.15 1 g 1015.40 Shipped in dry ice	1-DEOXY-1-NTRO-6-SORBITOL D 3526 [14199-88-3] C <sub>6</sub> H <sub>13</sub> NO <sub>7</sub> FW 211.2
	2'-DEOXYGUANOSINE 99-100% See also: Tissue Culture Media and Reagents Page 1758	25 mg 100 mg 250 mg 1 g	11.60 32.15 64.15 177.95	*	FW 507.2 (for free acid) R: 36/37/38 S: 26-36-22 EOXYGUANYLIC ACID (5')		DEOXYNIVALENOL  D 0156  (Vomitoxin; 3α,7α,15-Trihydroxy- 12,13-epoxytrichothec-9-en-8-one WARNING: Extremely hazardous! E
	[96]-07-9] C <sub>10</sub> H <sub>13</sub> N <sub>5</sub> O <sub>4</sub> FW 267.2 2'-DEOXYGUANOSINE-8-14C		592.80	D 0770	See: 2'-Deoxyguanosine 5'-Monop Page 356 '-DEOXYGUANYLYL(3'→5')- 2'-DEOXYGUANOSINE	1 mg 52.15 4 mg 173.70	risks and familiar with safety proci use this product.  Also available as part of a kit. See: I
	See: Radiochemicals Section Po	nge 21 <i>27</i> 5 mg	36.25	-o-c	Sodium Salt Minimum 98% [113753109] C <sub>20</sub> H <sub>24</sub> N <sub>10</sub> O <sub>10</sub> PNa		R: 26/27/28-36/37/38 S: 45-36 1-DEOXYNOJIRIMYCIN D 9305 Hydrochloride
D 7285	[3608-58-0] C <sub>10</sub> H <sub>13</sub> N <sub>5</sub> O <sub>4</sub> FW 267.2	10 mg	60.30	D 7514	-DEOXY-p-ribo-HEXOPYRANOSE [20789-85-9] C <sub>6</sub> H <sub>12</sub> O <sub>5</sub> FW 164.2	25 mg 10.95 100 mg 28.40 500 mg 101.60	A competitive inhibitor of glucosidase I and II <sup>1,2</sup> . Recently found to inhibit pig kidney trehalase Ref.: 1. Neverova, I., et al., Anal. E
D 9250	2'-DEOXYGUANOSINE 5'-DI- PHOSPHATE Sodium Salt Approx. 97%	100 mg 1		2	1-DEOXY-17-HYDROXYCORTICO See: Reichstein's Substance S Pag -DEOXY-20-HYDROXY- ECDYSONE	e 970 500 μg 40.65	190 (1994). 2. Yamashita, Y., et al., J. Virol., <b>61</b> 3. Kyosseva, S.V., et al., Arch. Biod <b>316</b> , 821 (1995).
	[102783-74-4] C <sub>10</sub> H <sub>15</sub> N <sub>5</sub> O <sub>10</sub> P <sub>2</sub> acid) R: 36/37/38 S: 26-36	FW 427.2 (for	free	2-8°C	(3β,14,20,22[R],25-Pentahydroxy- 5β-cholest-7-en-6-one) Minimum 80%		СН <sub>2</sub> ОН • НСІ
D 4147	'-DEOXYGUANOSINE 3'-MONO Ammonium Salt		 48.50	D 5287	[17942-08-4] C <sub>27</sub> H <sub>44</sub> O <sub>6</sub> FW 464. -DEOXYINOSINE Minimum 98%	100 mg 9.25 250 mg 15.30	но
<u>-o-c</u>	Approx. 98% [102783-49-3] C <sub>10</sub> H <sub>14</sub> N <sub>5</sub> O <sub>7</sub> P FW 347.2 (for free acid)		61.45 30.15	-orc	Essentially free of inosine. [890-38-0] C <sub>10</sub> H <sub>12</sub> N <sub>4</sub> O <sub>4</sub> FW 252.2 -DEOXYINOSINE	500 mg 25.40 1 g 46.95 5 g 193.95	[73285-50-4] C <sub>6</sub> H <sub>13</sub> NO <sub>4</sub> • HCl FV R: 36/38 S: 26-36 2-DEOXYNUCLEOSIDES and 2'-DEO
D 3264	<b>Sodium Salt Approx. 98%</b> [102814-03-9] C <sub>10</sub> H <sub>14</sub> N <sub>5</sub> O <sub>7</sub> P FW 347.2 (for free acid)	25 mg 1	48.50 61.45 30.15	D 0126 ⊡oc	5'-MONOPHOSPHATE Sodium Salt Synthetic	5 mg 7.65 100 mg 30.75 250 mg 61.35 1 g 170.35	5'-NUCLEOTIDES, Kits of See: Standards and Controls Section 3-DEOXYOCTULOSONIC ACID
2	R: 36/37/38 S: 26-36 <b>*-DEOXYGUANOSINE 5'-MONO</b>	PHOSPHATE			[14999-52-1] C <sub>10</sub> H <sub>13</sub> N <sub>4</sub> O <sub>7</sub> P FW 332.2 (for free acid) R: 36/37/38 S: 26-36 -DEOXYINOSINE	5 mg 14.90	See: 2-Keto-3-deoxyoctonate Page of p-erythro-DEOXYPENTOSE See: Deoxy-o-ribose Page 359
	(5'-Deoxyguanylic acid; d-GMP)	N		D 0758 	5'-TRIPHOSPHATE Sodium Salt Synthetic: 95-97%	25 mg 48.15 Shipped in dry ice	2-DEOXY-6-PHOSPHOGLUCONIC D 0376 ACID Sodium Salt
	O H <sub>2</sub> N N	h  }		<u>~</u>	[95648-77-4] C <sub>10</sub> H <sub>15</sub> N <sub>4</sub> O <sub>13</sub> P <sub>3</sub> FW acid) R: 36/37/38 S: 26-36 DEOXY-5'-S-ISOBUTYLTHIOADEI		Approx. 95% [102783-23-3] D 2681 - DEOXY-2-PHTHALIMIDO-
	· S			1- D 9160	See: 5'-S-IsobutyI-5'-Deoxyadenosin DEOXYMANNOJIRIMYCIN (1,5-Dideoxy-1,5-imino-p-mannitol)	ne Page 642 1 mg 25.45 5 mg 78.75	D 2681 3,4,6-TRI-O-ACETYL- α-D-GALACTOPYRANOSYL FLUORIDE Contains up to 10% β-anomer
D 9625	HO Free Acid 98-100% [902-04-5] C <sub>10</sub> H <sub>14</sub> N <sub>5</sub> O <sub>7</sub> P	500 mg 4	12.10 41.20		<b>Hydrochloride</b> [84444-906] C <sub>6</sub> H <sub>13</sub> NO <sub>4</sub> • HCI FW 199.6 R: 20/21/22 S: 36	10 mg 141.95	[17796656-2] C <sub>20</sub> H <sub>20</sub> NO <sub>9</sub> F FW 43 R: 36/37/38 S: 26-36 2-DEOXY-2-PHTHALIMIDO- D2806 3.4,6-TRI-0-ACETYL- 5
	FW 347.2 Sodium Salt 98-100%	 100 mg	8.50	6-1	DEOXY-L-MANNOSE See: L-Rhamnose Page 973 DEOXY-N <sup>6</sup> -METHYLADENOSINE	77.0	FLUORIDE May contain up to 10% o
	Also available as part of a kit. See: Standards and Controls Section Page 2148	500 mg 2 - 1 g 4	14.05	5′- D 5011	See: N <sup>6</sup> -Methyl-2'-deoxyadenosine <i>P</i> DEOXY-5'-METHYL- THIOADENOSINE	25 mg 19.45 100 mg 52.50	R: 36/37/38 S: 26-36
	[52558-16-4] C <sub>10</sub> H <sub>14</sub> N <sub>5</sub> O <sub>7</sub> P FW acid) R: 36/37/38 S: 26-36	347.2 (for free		-o·c	[2457-80-9] C <sub>11</sub> H <sub>15</sub> N <sub>5</sub> O <sub>3</sub> S FW 297.3 R: 36/37/38 S: 26-36	250 mg 104.20 1 g 271.30	0 0501 Hydrochloride Witamin B <sub>6</sub> antagonist [148-51-6] C <sub>6</sub> H <sub>11</sub> NO <sub>2</sub> • HCI FW 189.
356	Shipping info			Lla	<u> </u>		· · · · · · · · · · · · · · · · · · ·

## ALPHA

RODUCT UMBER	03 ;	NUMBER		USS	ALMER .
FITC-PEROXIDASE	cein Isothiocyanate labeled		AVIN ADENINE DINUCLEOTIDE (FAD) Disodium Salt Minimum 94% Orange powder.	10 mg 8.15 25 mg 11.35 100 mg 27.05 250 mg 54.10 500 mg 80.35	FLAVONE  F2003 (2-Phenyl-4H-1-benzo  Crystalline  [525-82-6] C <sub>15</sub> H <sub>10</sub> O <sub>3</sub> R: 36/37/38 S: 26-
FITC - PROTEIN A See: Protein A- FITC listed Page 947	d under Protein A - Soluble	-	HN N CH <sub>3</sub> CH <sub>3</sub>	1 g 144.80	FLAVORIDIN See: Tissue Culture M FLAZO ORANGE F2007 (1-15-Chloro-2-hydroxy
FIXATIVE SOLUTION See: Ethanol Fixative Page 45	2		H-C-OH H-C-OH H-C-OH O CH_O-P-O-P-OCH_OND OND OND	12 N	2-naphthol) Approx. 98% [35694-7] C <sub>16</sub> H <sub>11</sub> C FLECAINIDE
Fixing solution, Neutra  Fixing solution  Fixing solution	500 ml 66.3 1 liter 119.4		Also available as part of a kit. See: Standards and Controls Sect See also: Tissue Culture Media an Page 1759	d Reagents	F6777 (N-12-Piperidylmethyl)- [2,2,2-trifluoroethoxylt Acetate Salt Class I antiarrythmic al Ref.: 1. Roden, D.M. a Med., 315, 36 (1986). 2. Somani, P., Clin. Ph.
<ul> <li>17.5% (w/v) 5-sulfosalicy</li> <li>The working solution is upolyacrylamide and agar</li> <li>Suitable for PAGE, SDS-F</li> </ul>	ylic acid. Iseful for fixing proteins in Pose gels prior to staining. PAGE, and IEF systems.		[146-14-5] C <sub>27</sub> H <sub>31</sub> N <sub>9</sub> O <sub>15</sub> P <sub>2</sub> Na <sub>2</sub> F LAVIN ADENINE DINUCLEOTIDE See: Affinity Chromatography Med LAVIN MONONUCLEOTIDE	-AGAROSE	(1980). [54143-56-5] C <sub>17</sub> H <sub>20</sub> F R: 23/24/25-36/37/38 FLORISIL
X-Pro peptide bonds <sup>1</sup> (i.e isomerase) in synthetic s EK bioding protein chara	100 µg 106.7 cis-trans isomerization of a., a peptidyl prolyl substrates. reterized by binding to, and	*	(FMN; Riboflavin 5'-phosphate)	CH <sub>3</sub>	Magnesium silicate, ac The PR grade is suitable analysis.   F9760   Mesh: 16-30   Act. Temp. 1,250°F. [1343-88-0]   R: 36/37/38   S: 26-36   F3754   Mesh: 30-60   Act. Temp. 1,200°F. [1343-88-0]
43, 1101 (1984). 2. Handschumacher, R.E (1984). [13]] 44-19-9]	al., Biomed. Biochim. Acta, E., et al., Science, 226, 544  uct containing Metronidazole	F 8399	H - C - OH   CH <sub>2</sub> OH   (130-40-5)   C <sub>17</sub> H <sub>21</sub> N <sub>4</sub> O <sub>9</sub> P FW 4!   Sodium Salt   Approx. 95% (HPLC)   Prepared by the enzymatic	56.3 (for free acid) 1 mg 24.15 5 mg 80.30 10 mg 133.80	R: 36/37/38 S: 26-36  19127 Mesh: 60-100/PR Act. Temp. 1,250°F. [1343-88-0] R: 36/37/38 S: 26-36  Mesh: 60-100 Mesh: 60-100 Act. Temp. 1,200°F. [1343-88-0] R: 36/37/38 S: 26-36
(M 3761) Page 758  FLASHLIGHTS		F 2253	hydrolysis of flavin adenine dinucleotide. Sodium Salt; Synthetic Approx. 80% (HPLC) Riboflavin Content: Less than	10 mg 13.10 25 mg 25.40 100 mg 63.00	## 1752 Mesh: 100-200 Act. Temp. 1,200°F. ## 1343-88-01 ## 1343-88-01 ## 15638 S: 36
See: Techware Section	50 g 20. w 11) <b>50%</b>	90	O.3% A further purification of F 6750 to Biologically active in the growth of Strain 7469).  Also available as part of a kit. See: Standards and Controls Sec See also: Tissue Culture Media al Page 1759	o reduce riboflaving of L. casei (ATCC) ction Page 2148 and Reagents	1343-880   1343-880
FLAVIANIC ACID (2,4-Dinitro-1-naphthol-		F 6750	Sodium Salt Riboflavin Content: 73.0-79.0% Free Riboflavin: ≤6.0% Riboflavin Diphosphates: ≤6.0%	5 g 10.85 10 g 17.00 25 g 27.01 (as 100 g 86.85	PLOW CYTOMETRY COMPI See: Immunochemicals Post PUDARABINE des-PHOSPI See: 2-Fluoroadenine 9-β-D 10ge 483
F6500 Free Acid  [483-84-1] C₁₀H₀N₂O₃  R: 34-40 S: 26-36/37	25 g 27. S FW 314.2 7/39-22		riboflavin) Non-profit institutions may reque package GRATIS as often as nec gratis package per order.  FLAVIN MONONUCLEOTIDE, Ele		PUDROCORTISONE  100-Fluoro-11β,17α,21-trih  20-Fluorohydrocortisone; 20-Fluorocytisone; 20-Fluorocytisone; 20-Fluorocytisone; 20-Fluorocytisone;
F 7754 (C.I. 10316; Acid Yellor Naphthol Yellow S) Disodium Salt Dye content: Approx.	100 g 25	10	Reagent See: Electrophoresis Reagents F FLAVIN MONONUCLEOTIDE-AG See under: Affinity Chromatograf	page 1970 and	Etilosocortisone; etilosocortisol; 4-Pregnene itiol 3,20-dione) Approx. 98% [12:731-1] C <sub>21</sub> H <sub>29</sub> FO <sub>5</sub> FW

## **ALPH**

PRODUCT NUMBER	US \$	PRODUCT NUMBER	US\$ NUMBER	
	NHIBIN 1 vial 172.35 From Porcine Ovaries Follicle-stimulating hormone-suppressing protein 2,000 l.U. per vial Bioassay not run by Sigma. [57285-09-3]	l 2753	INOSINE 3':5'-CYCLIC	OSINE 5'-TRIPHOSP Periodate Oxidized (Inosine 5'-triphospha Sodium Salt: Minim [105208-87-5] C <sub>10</sub> hacid)
	R: 60 S: 45-36/37/39 INHIBIN-LIKE PEPTIDE	I 4375	5 Sodium Salt 100 mg 26.10 Approx, 97% 500 mg 113.05	OSINIC ACID See: Inosine 5'-Monor
	See: Bioactive Peptides Page 1218  INOCULATING LOOP/NEEDLE See: Techware Section Page 2325		Prepared from muscle or bacterial 1 g 203.75 ADP. [81012-88-6] C <sub>10</sub> H <sub>14</sub> N <sub>4</sub> O <sub>11</sub> P <sub>2</sub> FW 428.2 (for free acid)	i-INOSITOL (1,2,3,4,5/6-Hexahyd hexane) Approx. 95% [488-58-4] C <sub>6</sub> H <sub>12</sub> O <sub>6</sub>
	INORGANIC PYROPHOSPHATASE (Pyrophosphate phosphohydrolase; EC 3.6.1.1) Unit Definition: One unit will liberate 1.0 µmole of inorganic orthophosphate per min at pH 7.2 at 25°C, unless otherwise indicated. [9024-82-2]	17628	INOSINE, 2',3'-ISOPROPYLIDENE Derivative See: 2',3'-0-Isopropylideneinosine Page 647  INOSINE 3'-MONOPHOSPHATE 1 mg 9.90  B Sodium Salt 10 mg 50.65  Approx. 99%	ro-INOSITOL (1,2,3,5/4,6-Hexahyd hexane; meso-Inositol Minimum 99% See also: Tissue Cultu Reagents Page 1761 (87.89-8) C <sub>6</sub> H <sub>12</sub> O <sub>6</sub>
1891  -0°C	From Bakers Yeast 1 vial 38.35 HPLC purified, lyophilized Activity: 500-1,500 units per mg protein (BCA). Prepared from I 1643; essentially salt-free Minimum 90% (reversed phase HPLC)	-	[97259-68-2] C <sub>10</sub> H <sub>13</sub> N <sub>4</sub> O <sub>8</sub> P FW 348.2 my   INOSINE 5'-MONOPHOSPHATE (Inosinic Acid; IMP; I-5'-P) sc)	o-INOSITOL-[2-3H] See: Radiochemicals S IIIo-INOSITOL
I 1643 -⊙°c	Vial contains 100 µg protein.  From Bakers Yeast 100 units 22.70 Lyophilized powder 500 units 65.10		Grade V: 98-100%   1 g   14.45   m   f	DTLET; 1,3,5/2,4,6-h nydroxycyclohexane) [488-59-5] C <sub>6</sub> H <sub>12</sub> O <sub>6</sub> o-INOSITOL, 2,2'-AI
1 2267 2-8°C	containing 85% buffer 1,000 units 119.90 salts.  Activity: 500-1,000 units per mg protein (E½%).  From E. coli 100 units 21.70 Minimum 60% (SDS-PAGE) 500 units 62.25 Lyophilized powder containing Tris buffer salts.  Activity: Minimum 1,000 units per mg protein (E½%)		HN N N	2-C-HYDROXYMETH 2-C-Methylene-myo-in Approx. 99% Reported to be a compositol. Ref.: Posternak, T., "1 rancisco, Ca. (1965). 4068-87-5] C <sub>7</sub> H <sub>12</sub> O <sub>6</sub>
1 2891 2-8°C	at pH 9.0 at 25°C.  Thermostable Enzyme 50 units 25.96 from Bacillus 250 units 102.24 stearothermophilus 1,000 units 306.86 Lyophilized Activity: 15-25 units per mg protein (Biuret) at pH 9.0 at 50°C.	; <b>]</b>	HO OH  [131-99-7] C <sub>10</sub> H <sub>13</sub> N <sub>4</sub> O <sub>8</sub> P FW 348.2  5 Disodium Salt 1 g 6.40 Grade III: 98-100% 5 g 13.25 From Yeast 10 g 22.15 Crystalline 25 g 38.10	yo-INOSITOL 1,4-bi PHATE Potassium Salt upprox. 98% (TLC) rom Bovine Brain L- Monophosphate tef.: Emilsson, A. and 159, 3111 (1984).
	INORGANIC PYROPHOSPHATES See: Pyrophosphates Page 875	1 450	[4691-650] C <sub>10</sub> H <sub>11</sub> N <sub>4</sub> O <sub>8</sub> PNa <sub>2</sub> 100 g 134.65 [ FW 392.2 a	1 <i>03476-30-8</i> ]
	(Hypoxanthine 9-p-ribofuranoside) We also offer: 2',3'-0-Isopropylideneinosine Page 647	2-8°C	Sigma Grade: 99-100%	HOSPHATE mmonium Salt ot assayed by Sigma. 106358-02-5] C <sub>6</sub> H <sub>14</sub>
l 1024 RT	[5863-9] C <sub>10</sub> H <sub>12</sub> N <sub>4</sub> O <sub>5</sub> FW 268.2  SIgmaUltra 25 g 58.8  Minimum 99% 100 g 214.0  Residue upon ignition: <0.1%  Solubility (0.5 M in water, 20°C): complete, colorless		INOSINE MONOPHOSPHATE, CYCLIC See: Inosine 3':5'-Cyclic Monophosphate Page 628 INOSINE 5'-MONOPHOSPHATE, 25 mg 36.10	yo-INOSITOL 4,5-bl: HOSPHATE mmonlum Salt ot assayed by Sigma. 59256-54-8] C <sub>6</sub> H <sub>14</sub> O
	Insoluble matter: <0.2% CI: <0.05% SO <sub>4</sub> : <0.05% AI: <0.0005 AI: <0.0005% CI: <0.0005% P: <0.0005 CU: <0.0005% Pb: <0.001	% -0°C	(Inosine 5'-monophosphate-2',3'-dialdehyde)  Sodium Salt: Minimum 80%  Balance Inorganic salts [112898-40-5] C <sub>10</sub> H <sub>11</sub> N <sub>4</sub> O <sub>8</sub> P FW 346.2 (for free acid)	myo-INOSITOL 5,6- HOSPHATE yclohexylammoniur ot assayed by Sigma 142507-73-1} C <sub>6</sub> H <sub>18</sub> ! cid)
l 4125 RT	Fe: <0.0005% Zn: <0.0005 K: <0.005% 1 g 6.4 5 g 11.0 10 g 18.3 25 g 29.4 100 g 107.0	5 5 1 087 5 0 •	See: Inosine Mono, Di, or Tri-phosphate	yo-INOSITOL 1,2-C yclohexylammoniur pprox. 98% (TLC) 26/38-12-91 C <sub>6</sub> H <sub>11</sub> O <sub>1</sub> 10-36/37/38 S: 16 mg per ml solution in
	INOSINE-8- <sup>14</sup> C See: Radiochemicals Section Page 2133	*   *	[35908-31-7] C10H12N4O14P3Na3	ater (3:1).
628	◆ Shipping information - page 5		How to use catalog - page 2.	To place a

носн<sub>2</sub>

[54-42-2] C<sub>9</sub>H<sub>11</sub> R: 45-46-61-43 36/37/39-22

5-IODO-2'-DEOXYI 5'-MONOPHOSF

Sodium Salt

Approx. 98% [103404-69-9] acid)

5-IODO-2,4-DIMET PYRIMIDINE

5-HODO-1,3-DIMETI 99% (HPLC) [40738-83-8] C<sub>6</sub> R: 63-20/21/22-4

**IODOETHANE** 

(Ethyl iodide) d = 1.95 g/ml Colorless to faint y Stabilized with 0.3 [75-03-6]  $C_2H_5$ | R: 23/24/25-63-4

2-IODOETHANOL

d = 2.2 g/ml [624-76-0] C₂H₅I(

R: 46-23/24/25-3

N-(2-IODOETHYL)TF ACETAMIDE [67680-56-2] C<sub>4</sub>1-R: 36/37/38 S: 2

> (Triiodomethane) Yellow crystals. [75-47-8] CHI<sub>3</sub> FN R: 20/21/22-36/3;

> Trademark of Pierce 1,3,4,6-Tetrachloro See: Page 1045

7-IODO-8-HYDROXYI 5-SULFONIC ACID

IODOFORM

IODO-GEN

Minimum 98% ( Ref.: Kundu, N.G. Trans., 1991, 10. [52522-99-3] C<sub>ε</sub>

19878

-o·c

18766

19020

14259

12507

2-8°C

1 2257 24°C

11131

3753

5-IODO-2'-DEOXY 17125 (IDU; Idoxuridine Minimum 99%

PRODUCT NUMBER			us \$	PRODUCT NUMBER			US \$
(C	ontinuation of) DOACETIC ACID Free Acid Approx. 99% Yellow powder. May form hazy solution in water.	10 g 25 g 100 g	12.85 25.60 85.75	1 7626 aT	D-IODOBENZOIC ACID (2-Iodobenzoic acid) Light yellow crystals. [88-67-5] C.H <sub>3</sub> [0 <sub>2</sub> FW 248.0 R: 20/21/22-42/43-40 S: 26-36-22	5 g 25 g 100 g 250 g	4.05 11.80 32.05 70.55
I 1014 ⊡©	[64-6-7] C <sub>2</sub> H <sub>3</sub>  O <sub>2</sub> FW 185.9 R: 25-35 S: 22-36/37/39-45 Lthhium Salt Minimum 97% (titration) [65749-30-6] C <sub>2</sub> H <sub>2</sub>  O <sub>2</sub> Li FW 191.	_	22.70	1 6500 2-8°C ◆	o-IODOBENZOTRIFLUORIDE (2-[Trifluoromethyl]iodobenzene) d = 1.90 g/ml [444-29-1] C <sub>3</sub> H <sub>4</sub> F <sub>3</sub>   FW 272.0 R: 34 S: 26-27-36/37/39	1 g 5 g 25 g	4.40 8.50 24.50
9148  -∞c   ◆	R: 20/21/22-36/37/38 S: 26-36  Sodium Salt SigmaUltra Approx. 99% Solubility (0.5 M in water, 20°C): co Insoluble matter: <0.1% SO <sub>4</sub> : <0.05% AI: <0.0005% Ca: <0.0005% Cu: <0.001%	K: Mg: < NH₄* P: <	28.90 96.05 lorless < 0.02% 0.0005% < 0.05% 0.0005%	1 9890 2-8°C	m-IODOBENZYLGUANIDINE (MIBG) Hemisulfate Salt Antitumor agent which inhibits ADP Ref.: 1. Smets, L.A., et al., Cancer Pharmacol., 21, 9 (1988). 2. Loesberg, C., et al., Biochim. Bi 1037, 92 (1990). [80663-95-2] C <sub>8</sub> H <sub>10</sub> IN <sub>3</sub> • 1/2H <sub>2</sub> S0	ribosylation Chemothe	r. .,
12512	Fe: <0.0005% [305-53-3] C <sub>2</sub> H <sub>2</sub> IO <sub>2</sub> Na FW 207.9 R: 23/24/25 S: 45-26-36/37/39 Sodium Salt	22 25 g	0.0005%  45.75 126.65		IODOCHLOROHYDROXYQUINOLIN See: 5-Chloro-7-iodo-8-hydroxyquir 3β-IODO-5-CHOLESTENE See: Cholesteryl lodide Page 284	E oline Page	270
-o·c ◆	Approx. 99% [305-53-3] C <sub>2</sub> H <sub>2</sub> IO <sub>2</sub> Na FW 207.9 R: 23/24/25 S: 45-26-36/37/39				19-IODO-5-CHOLESTEN-3β-OL 3-A See: 19-Iodocholesterol 3-Acetate	ACETATE	
। । 9760 - <u>ञ</u> ्च	DDACETIC ACID N-HYDROXY- SUCCINIMIDE ESTER Reagent for cross-linking proteins. Ref.: 1. J. Immun. Meth., 24, 321 (1978). 2. Eur. J. Biochem., 140, 63 (198- [39028-278] C <sub>6</sub> H <sub>6</sub> INO <sub>4</sub> FW 283 R: 36/37/38 S: 26-36	4).	14.10 53.85 96.45		19-IODOCHOLESTEROL 3-ACETAT (5-Cholesten-19-iodo-3β-ol 3-acetate; 19-iodo-5-cholesten-3β Approx. 95% Crystalline [4561-904] C <sub>29</sub> H <sub>46</sub> IO <sub>2</sub> FW 553.	E 1 mg	44.95 e)
I 3507 2-8°C	ODOACETIC ANHYDRIDE [54907-61-8] C₄H₄I₂O₃ FW 353.9 R: 34-23/24/25 S: 26-27- 36/37/39	100 mg 250 mg 1 g		16128	5-IODOCYTIDINE (4-Amino-2-hydroxy-5-iodo- 1β-o-ribofuranosylpyrimidine) Crystalline [1147-23-5] C <sub>9</sub> H <sub>12</sub> IN <sub>3</sub> O <sub>5</sub> FW 369	25 mg 100 mg	27.75 73.40
8879   <u> </u> <del> </del>	N-IODOACETYL-N'-(5-SULFO- 1-NAPHTHYL)ETHYLENE- DIAMINE (1,5-I-AEDANS) Minimum 80% (HPLC) Yellow crystals.	1 g	16.15 89.40	I 5628	5-IODOCYTIDINE 5'-TRIPHOS- PHATE Sodium Salt Approx. 95% [118357-27-0] C <sub>9</sub> H <sub>15</sub> IN <sub>3</sub> O <sub>14</sub> P <sub>3</sub> F acid)	5 mg 25 mg W 609.1 (fo	
   9004  RT	[36930-63-9] C <sub>14</sub> H <sub>15</sub> IN <sub>2</sub> O <sub>4</sub> S FW N-IODOACETYL-N'-(8-SULFO- 1-NAPHTHYL)ETHYLENE- DIAMINE (1,8-I-AEDANS)	25 mg	19.60	1 6875	5-IODOCYTOSINE (4-Amino-2-hydroxy-5-iodo- pyrimidine) Crystalline [1122-44-7] C <sub>4</sub> H <sub>4</sub> IN <sub>3</sub> O FW 237	100 mg 500 mg 1 g 5 g	23.05 37.65 121.35
1 1757	Yellow crystals. [36930-64-0] C <sub>14</sub> H <sub>15</sub> IN <sub>2</sub> O <sub>4</sub> S FW  19-IODO-5-ANDROSTENE-3β-OL- 17-ONE 3-ACETATE	5 mg	148.00	1 5883 -oc	5'-IODO-5'-DEOXYADENOSINE Minimum 95% Crystalline [4099-81-4] C <sub>10</sub> H <sub>12</sub> IN <sub>5</sub> O <sub>3</sub> FW 3	100 mg	40.75
	[82341-96-6] C <sub>21</sub> H <sub>29</sub> IO <sub>3</sub> FW 450 5-10D0ANTHRANILIC ACID See: 2-Amino-5-lodobenzoic Acid	Page 111		17000  -0°C	5-IODO-2'-DEOXYCYTIDINE ) Crystalline [611-53-0] C <sub>9</sub> H <sub>12</sub> IN <sub>3</sub> O <sub>4</sub> FW 353.1	100 mg 1 g 5 g	15.20 86.35 341.80
	4-IODOANTIPYRENE-N-METHYL-1 See: Radiochemicals Section Page p-IODOBENZENESULFONYL CHLC	2133		I 826:	5-10DO-2'-DEOXYCYTIDINE	1 mg 5 mg	20.75 68.35
1 4759	See: Pipsyl Chloride Page 903  4-IODOBENZOIC ACID [619-58-9] C <sub>1</sub> H <sub>3</sub> IO <sub>2</sub> FW 248.0 R: 36/37/38 S: 26-36	5	g 14.3	- 1	Approx. 95% [31747-59-8] C <sub>9</sub> H <sub>15</sub> IN <sub>3</sub> O <sub>13</sub> P <sub>3</sub> Flacid) R: 23/24/25-36/37/38 S: 26-3		r free

(lodoxyquinolinesulfc [547-91-1] C<sub>9</sub>H<sub>6</sub>INt R: 34 S: 26-28-27 To plac

(Continua IONOPH Amma Cockt

> Bariui (V 163 oxybis [964] Cadm

3,6-di [7348

Calci

(ETH ) [5880

Calcii Cocki R: 36, Calcii

Cock R: 36,

Calci

N,N,N [742c R: 37

Calci Cock R: 10 Calci

Cock R: 11

Calci See: (

Calci (ETH N',N'-[126.

Carb (ETH 4-trifl In 0.5 [129 R: 11

Carb

4-trifl In O.!

Cart (ETH

4-trif 5 ml [129 R: 1] Chrc (ETH

5-oc [125

(ETH 5-[4phen [134]

11272 EFC 11397 EFC

1522 2-5°C

1647

1772 28°C

2-8°C

12022

2-8°C

12147 2-8°C

12272

12522

1 2647 2-8℃

12772

2-8°C

Œ.

12897 ≥8℃

13147

1 3272 2-FC

2-8°C

PRODUCT NUMBER			US \$	PRODUC' NUMBER			US
I 7509 -o⁺c	p-IODO-p-PHENYLALANINE (2-Amino-3-[4-iodophenyl]-propano acid) [62561-75-5] C <sub>9</sub> H <sub>10</sub> NO <sub>2</sub> FW 29	ic 1.1	127.70	I 8250 -⊙°C	3-IODO-t-TYROSINE (3-Monoiodo-t-tyrosine) Crystalline Contains approx. 5% tyrosine. (70-78-0) CaHiolNO <sub>3</sub> FW 307.1	1 g 5 g 25 g	20.1 67.0 223.4
I 4628 RT	p-IODO-oL-PHENYLALANINE (2-Amino-3-[4-iodophenylpropanoiacid) Crystalline	100 mg 500 mg 1 g 5 g			L-m-IODOTYROSINE See: 3-lodo-t-tyrosine Page 638		
I 8757	[14173:41-2] C <sub>9</sub> H <sub>10</sub> INO <sub>2</sub> FW 291.1 <b>p-IODO-L-PHENYLALANINE</b> (2-Amino-3-[4-iodophenyl]- propanoic acid) [24250:85:9] C <sub>9</sub> H <sub>10</sub> INO <sub>2</sub>	500 mg 1 g 5 g	26.80 48.20 190.70	I 5016	5-IODOURACIL (2,4-Dihydroxy-5-iodopyrimidine) Minimum 98% [696-07-1] C <sub>4</sub> H <sub>3</sub> IN <sub>2</sub> O <sub>2</sub> FW 238.0 R: 46-20/21/22-36/37/38 S: 45- 26-36/37/39-22	1 g 5 g 10 g 25 g	5. 15. 25. 56.
2146	FW 291.1  4-(o-IODOPHENYL)BUTYRIC ACID [159002-37-6]  4-(o-IODOPHENYL)BUTYRIC ACID		Inquire	I 7500	furanosylpyrimidine) Crystalline	250 mg	13.
1 5634 2-8°C	[27913:58:2] C <sub>10</sub> H <sub>11</sub> IO <sub>2</sub> FW 290.1 <b>2-(p-IODOPHENYL)-3-p-NITROPHE</b>		15.75 27.60 105.75	I 8378	[1024:99:3] C <sub>9</sub> H <sub>11</sub> IN <sub>2</sub> O <sub>6</sub> FW 370.  5-IODOURIDINE 5'-MONO-PHOSPHATE Sodium Salt	1 5 mg	15.
1 0256	5-PHENYLTETRAZOLIUM CHLOI See: p-lodonitrotetrazolium Violet (I IODOPLATINATE SPRAY REAGENT 0.15% Potassium chloroplatinate	NT) Page 6	23.00		Approx. 98% Crystalline [103404-82-6] C <sub>9</sub> H <sub>12</sub> IN <sub>2</sub> O <sub>9</sub> P FW 4 acid)	150.1 (for	free
2-8°C	and 3% potassium iodide in dilute he For use in the detection of alkaloids organic nitrogen compounds. R: 40-36/37/38 S: 26-36			I 3012 ⊡⊙©	5-IODOURIDINE 5'-TRIPHOSPHATE Sodium Salt Approx. 95% [73431-55-7] C <sub>9</sub> H <sub>14</sub> IN <sub>2</sub> O <sub>15</sub> P <sub>3</sub> FW 6	1 mg 10 mg	18. 101.
I 9882 RT	1-IODOPROPANE Approx. 99% Stabilized with copper. d = 1.74 g/ml	100 ml	30.50		acid) R: 23/24/25-36/37/38-42/43-40 36/37/39	S: 45-26-	
	[107-08-4] C <sub>3</sub> H <sub>7</sub> I FW 170.0 R: 10-45-36/37/38 S: 16-45-26- <b>2-IODOPROPANE</b>	36/37/39 100 g	15.55		IODOXYQUINOLINESULFONIC ACID See: 7-lodo-8-hydroxyquinoline-5-sul Page 637	fonic Acid	
I 0133	Stabilized with copper. d = 1.70 g/ml Possible carcinogen. (75·30-9) C <sub>3</sub> H <sub>3</sub> I FW 170.0 R: 10-20/21/22-36/37/38-40 S:			1 0634 2-8°C	IONOMYCIN Calcium Salt From Streptomyces conglobatus Ca <sup>2+</sup> ionophore that is more effectiv a mobile ion carrier for Ca <sup>2+</sup> ; non-flu study Ca <sup>2+</sup> transport across biologic induces apoptotic neuronal degener	e than A23 orescent; al membra	187 used
1 7875 2-8℃	6-IODOPURINE Crystalline [2545-26-8] C <sub>5</sub> H <sub>3</sub> IN <sub>4</sub> FW 246.0 4-IODOPYRAZOLE	10 g	64.20		embryonic cortical neurons  Ref.: Toeplitz, B.K., et al., J. Am. Cl 3344 (1979).  [56092-82-1] C <sub>41</sub> H <sub>70</sub> O <sub>9</sub> Ca FW 74	nem. Soc.,	101
I 6003	Crystalline [3469690] C <sub>3</sub> H <sub>3</sub> IN <sub>2</sub> FW 194.0 R: 42/43-40 S: 26-36-22			1 3384	R: 22 S: 36 α-IONONE	100 g	25.
1 8000 ( RT	o-IODOSOBENZOIC ACID (2-lodosobenzoic acid) Crystalline [304-91-6] C <sub>7</sub> H <sub>5</sub> IO <sub>3</sub> FW 264.0	1 g 5 g	8.15 32.20	2-8°C	d = 0.93 g/ml {127-41-3} C <sub>13</sub> H <sub>20</sub> O FW 192.3 R: 42/43 S: 36		
3761 -20°C	R: 36/37/38 S: 26-36 9(10)-IODOSTEARIC ACID 98+% [112966-11-7] C <sub>18</sub> H <sub>35</sub> IO <sub>2</sub> FW 41	500 mg 0.4	43.25	I 6381 2-8°C	β-IONONE (4-[2,6,6-Trimethyl-1-cyclohexen- 1-yl]-3-buten-2-one) Minimum 95% (GC) d = 0.95 g/ml	25 ml 100 ml	13. 22.
1 3886 -20°C	9(10)-IODOSTEARIC ACID METHYL ESTER Approx. 97% [112897-95-7] C <sub>19</sub> H <sub>37</sub> IO <sub>2</sub>	250 mg 500 mg 1 g	31.65 56.30 100.90		[79-77-6] C <sub>13</sub> H <sub>20</sub> O FW 192.3 R: 42/43 S: 36 IONOPHORES		
	FW 424.4 N-IODOSUCCINIMIDE	1 g 5 g	9.65 25.40		FlukaBrand Selectophore®  Selectophore ionophores and cockta for production of reliable and accura electrodes.		
7142 2-8°C	Minimum 95% Orange powder.	10 g	45.70		.,	0.1 ml	

#### Attachment 3

	Peptide sequence	Reference
1	RLEYYENEKK	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.
2	KRGEEELSNYICMGGK	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.
3	KKVSIEEYTEMMPAK	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.
4	KKHTDDGYMPMSPGVA	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.
5	RKGNGDGYMPMSPKSV	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.
6	KKRVDPNGYMMMSPSGS	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.
7	KKKLPATGDYMNMSPVGD	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992. Hubbard, EMBO J 16:5573-5581, 1997.
8	KKGSEEYMNMDLGPGR	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.
9	KKSRGDYMTMQIG	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.
10	KPRNSYVDTSPVAPK	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.
11	KKSRGNYMTMQIG	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.
12	KKSRGDYITMQIG	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.
13	KKSRGDYTTMQIG	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.

	Peptide sequence	Reference
14	KKSRGDY(Nle <sup>1</sup> )TMQIG	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.
15	KKSRGDYMTTQIG	Shoelson et al., Proc. Natl. Acad. Sci. USA 89:2027-2031, 1992.
16	TRDIYETDYYRK	Stadmauer et al., J Biol. Chem. 261:10000-10005, 1996.
17	LFASSNPEYLSARR	Stadmauer et al., J Biol. Chem. 261:10000-10005, 1996.
18	KRSYEEHIPYTHMNGGK	Stadmauer et al., J Biol. Chem. 261:10000-10005, 1996.
19	SRYMEDSTYYKASKG	Baron et al., J. Biol. Chem. 273:7162-7168, 1998.

<sup>&</sup>lt;sup>1</sup> Norleucine

#### Attachment 4

	Peptide sequence	Kinase	Reference
20	PLSRTLSVSS	PKC <sup>2</sup>	Kwon et al., J. Biol. Chem. 269:4839-4844, 1994.
21	PLSRTLSV	PKC	Kwon et al., J. Biol. Chem. 269:4839-4844, 1994.
22	PLSRTLS	PKC	Kwon et al., J. Biol. Chem. 269:4839-4844, 1994.
23	PLRRTLSVAA	PKC	Kwon et al., J. Biol. Chem. 269:4839-4844, 1994.
24	PLSRRLSVAA	PKC	Kwon et al., J. Biol. Chem. 269:4839-4844, 1994.
25	KKKKKRFSFKKAFKKLA- GFAFKKNK	PKC	Kwon et al., J. Biol. Chem. 269:4839-4844, 1994.
26	DEDADIYDEEDYDL	CK2 <sup>3</sup>	Marin et al., J. Biol. Chem. 274:29260-29265, 1999.
27	DEDADIYDEADYDL	CK2	Marin et al., J. Biol. Chem. 274:29260-29265, 1999.
28	DEDADIYDAEDYDL	CK2	Marin et al., J. Biol. Chem. 274:29260-29265, 1999.
29	DEDADIYAEEDYDL	CK2	Marin et al., J. Biol. Chem. 274:29260-29265, 1999.
30	DEDADDYDEEDYDL	CK2	Marin et al., J. Biol. Chem. 274:29260-29265, 1999.
31	DEDADISDEEDYDL	CK2	Marin et al., J. Biol. Chem. 274:29260-29265, 1999.

<sup>&</sup>lt;sup>2</sup> Protein Kinase C <sup>3</sup> Casein kinase-2

	Peptide sequence	Kinase	Reference
32	DEDADDSDEEDYDL	CK2	Marin et al., J. Biol. Chem. 274:29260-29265, 1999.
33	DEDADISAEEDYDL	CK2	Marin et al., J. Biol. Chem. 274:29260-29265, 1999.
34	DEDADISDEADYDL	CK2	Marin et al., J. Biol. Chem. 274:29260-29265, 1999.
35	RRREEEEESAAA	GRK2 <sup>4</sup>	Onorato et al., J. Biol. Chem. 270:21346-21353, 1995.
36	VSRSGLYRSPSMPENLNRP- RL	Chk1 <sup>5</sup>	Hutchins et al., FEBS Lett. 466:91-95, 2000.
37	LNRSRLYRSPSMPEKLDR- MPL	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
38	TPRRTLFRSLSCTVETPLA- NK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
39	YLRPNVSRSRSSGNAPPFL- RS	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
40	QDTPVVRRTQSMFLNST- RLGL	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
41	RLYRSPSMPEKLD	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
42	ALYRSPSMPEKLD	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
43	RAYRSPSMPEKLD	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
44	RLARSPSMPEKLD	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.

<sup>&</sup>lt;sup>4</sup> G protein coupled receptor kinase <sup>5</sup> Checkpoint kinase-1

	Peptide sequence	Kinase	Reference
45	RLYASPSMPEKLD	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
46	RLYRAPSMPEKLD	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
47	RLYRSASMPEKLD	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
48	RLYRSPAMPEKLD	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
49	RLYRSPSAPEKLD	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
50	RLYRSPSMAEKLD	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
51	RLYRSPSMPAKLD	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
52	RLYRSPSMPEALD	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
53	RLYRSPSMPEKAD	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
54	RLYRSPSMPEKLA	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
55	RLYRAPSMPEKLDRK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
56	RLARAASMAAALARK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
57	RVARAASMAAALARK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
58	RMARAASMAAALARK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.

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	Peptide sequence	Kinase	Reference
59	RRARAASMAAALARK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
60	RIARAASMAAALARK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
61	RAARAASMAAALARM	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
62	RLAKAASMAAALARK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
63	RLAAAASMAAALARK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
64	RLARAASMAAAAARK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
65	RLARAASMAAAIARK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
66	RLARAASMAAAVARK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
67	RLARAASMAAAALRK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
68	RLARAASMAALAARK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
69	RLARAASAAAAAARK	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
70	RKRLARAASMAAALA	Chk1	Hutchins et al., FEBS Lett. 466:91-95, 2000.
71	SAVGFNEMEAPTTAYK	Lyn <sup>6</sup>	Yamanashi et al., Proc. Natl. Acad. Sci. USA 90:3631- 3635, 1993.

<sup>&</sup>lt;sup>6</sup> Cellular Lyn protein kinase

	Peptide sequence	Kinase	Reference
72	KKLIEDAGYAARG	c-Abl <sup>7</sup>	Till et al., J. Biol. Chem. 274:4995-5003, 1999.
73	KKLIEDAIYAARG	c-Abl	Till et al., J. Biol. Chem. 274:4995-5003, 1999.
74	KKLIEDALYAARG	c-Abl	Till et al., J. Biol. Chem. 274:4995-5003, 1999.
75	KKLIEDAHYAARG	c-Abl	Till et al., J. Biol. Chem. 274:4995-5003, 1999.
76	KKLIEDAAYAARG	c-Abl	Till et al., J. Biol. Chem. 274:4995-5003, 1999.
77	KKLIEDAKYAARG	c-Abl	Till et al., J. Biol. Chem. 274:4995-5003, 1999.
78	KKLIEDAQYAARG	c-Abl	Till et al., J. Biol. Chem. 274:4995-5003, 1999.
79	KKSRGDYMTMQIG	c-Abl, v-Abl <sup>8</sup> , v-Src <sup>9</sup>	Till et al., J. Biol. Chem. 274:4995-5003, 1999; Garcia et al., J. Biol. Chem. 268:25146-25151, 1993.
80	KKSRGDYITMQIG	c-Abl, v-Abl, v-Src	Till et al., J. Biol. Chem. 274:4995-5003, 1999; Garcia et al., J. Biol. Chem. 268:25146-25151, 1993.
81	KKSRGDY(Nle) <sup>10</sup> TMQIG	c-Abl, v-Abl, v-Src	Till et al., J. Biol. Chem. 274:4995-5003, 1999; Garcia et al., J. Biol. Chem. 268:25146-25151, 1993.

 <sup>&</sup>lt;sup>7</sup> Cellular Abl protein kinase
 <sup>8</sup> Viral Abl protein kinase
 <sup>9</sup> Viral Src protein kinase
 <sup>10</sup> Norleucine

	Peptide sequence	Kinase	Reference
82	KKSRGDYATMQIG	c-Abl	Till et al., J. Biol. Chem. 274:4995-5003, 1999.
83	KKSRGDYETMQIG	c-Abl	Till et al., J. Biol. Chem. 274:4995-5003, 1999.
84	KKSRGDYMTPQIG	c-Abl	Till et al., J. Biol. Chem. 274:4995-5003, 1999.
85	KKSRGDYMTTQIG	c-Abl, v-Abl, v-Src	Till et al., J. Biol. Chem. 274:4995-5003, 1999; Garcia et al., J. Biol. Chem. 268:25146-25151, 1993.
86	KKSRGDYMTAQIG	c-Abl	Till et al., J. Biol. Chem. 274:4995-5003, 1999.
87	KKSRGDYMTEQIG	c-Abl	Till et al., J. Biol. Chem. 274:4995-5003, 1999.
88	KKHTDDGYMPMSPGVA	v-Src, v-Abl	Garcia et al., J. Biol. Chem. 268:25146-25151, 1993.
89	RKGNGDGYMPMSPKSV	v-Src, v-Abl	Garcia et al., J. Biol. Chem. 268:25146-25151, 1993.
90	KKRVDPNGYMMMSPSGS	v-Src, v-Abl	Garcia et al., J. Biol. Chem. 268:25146-25151, 1993.
91	KKKLPATGDYMNMSP- VGD	v-Src, v-Abl	Garcia et al., J. Biol. Chem. 268:25146-25151, 1993.
92	KKGSEEYMNMDLGPGR	v-Src, v-Abl	Garcia et al., J. Biol. Chem. 268:25146-25151, 1993.
93	KKKEEEEEEYMPMEDL	v-Src, v-Abl	Garcia et al., J. Biol. Chem. 268:25146-25151, 1993.

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	Peptide sequence	Kinase	Reference
94	KKSRGNYMTMQIG	v-Src, v-Abl	Garcia et al., J. Biol. Chem. 268:25146-25151, 1993.
95	KKSRGDYTTMQIG	v-Src, v-Abl	Garcia et al., J. Biol. Chem. 268:25146-25151, 1993.
96	ADFGLARLIEDNEYTARG	c-Src <sup>11</sup> , Hck <sup>12</sup>	Silicia et al., J. Biol. Chem. 273:16756-16763, 1998.
97	AEEEIYGEFEAKKKK	c-Src, Hck	Silicia et al., J. Biol. Chem. 273:16756-16763, 1998.
98	AEEEAYGEAEAKKKK	c-Src, Hck	Silicia et al., J. Biol. Chem. 273:16756-16763, 1998.
99	AEVIYAAPFAKKKK	c-Src, Hck	Silicia et al., J. Biol. Chem. 273:16756-16763, 1998.
100	KVEKIGEGTYGVVYK	c-Src, Hck	Silicia et al., J. Biol. Chem. 273:16756-16763, 1998.
101	KVEKIGEGTYGVVKK	c-Src, Hck	Silicia et al., J. Biol. Chem. 273:16756-16763, 1998.
102	KVEKIGVGSYGVVKK	c-Src, Hck	Silicia et al., J. Biol. Chem. 273:16756-16763, 1998.

<sup>11</sup> Cellular Src protein kinase 12 Src-like protein kinase

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